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FILE 'HOME' ENTERED AT 16:21:28 ON 02 NOV 2001

=> file medline, uspat, dgene, hcaplus, embase

COST IN U.S. DOLLARS

SINCE FILE

ENTRY SESSION 0.30

TOTAL

FULL ESTIMATED COST

FILE 'MEDLINE' ENTERED AT 16:32:16 ON 02 NOV 2001

FILE 'USPATFULL' ENTERED AT 16:32:16 ON 02 NOV 2001 CA INDEXING COPYRIGHT (C) 2001 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'DGENE' ENTERED AT 16:32:16 ON 02 NOV 2001 COPYRIGHT (C) 2001 DERWENT INFORMATION LTD

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FILE 'EMBASE' ENTERED AT 16:32:16 ON 02 NOV 2001 COPYRIGHT (C) 2001 Elsevier Science B.V. All rights reserved.

=> s fluorescent dye

L1 21604 FLUORESCENT DYE

=> s 12 and tetracysteome

L2 NOT FOUND

The L-number entered could not be found. To see the definition of L-numbers, enter DISPLAY HISTORY at an arrow prompt (=>).

=> s l1 and tetracysteine

L2 0 L1 AND TETRACYSTEINE

=> s fluorescein arsenical helix binder

L3 3 FLUORESCEIN ARSENICAL HELIX BINDER

=> s protein tag

L4 212 PROTEIN TAG

=> s FLAsh

L5 148618 FLASH

=> s 15 and cysteine

L6 2081 L5 AND CYSTEINE

=> s 16 and 13

L7 0 L6 AND L3

=> s 16 and 14

L8 0 L6 AND L4

=> d 13 ti abs ibib tot

ANSWER 1 OF 3 USPATELL L3

Compositions and methods for assaying subcellular conditions and TI processes using energy transfer

The invention is provides compositions and methods for monitoring AB subcellular compartments such as organelles by energy transfer techniques that do not require specific intermolecular affinity binding events between energy transfer donor and energy transfer acceptor molecules. Provided are methods for assaying cellular membrane potential, including mitochondrial membrane potential, by energy transfer methodologies including fluorescence resonance energy transfer (FRET). Diagnostic and drug screening assays are also provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:142122 USPATFULL

Compositions and methods for assaying subcellular TITLE:

conditions and processes using energy transfer

Dykens, James A., Encinitas, CA, United States INVENTOR(S):

Veli.cedilla.elebi, Gonul, San Diego, CA, United

States

to

Ghosh, Soumitra S., San Diego, CA, United States

Mitokor, San Diego, CA, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE US 6280981 B1 20010828 PATENT INFORMATION: US 2000-514569 20000223 (9)

Division of Ser. No. US 1999-338122, filed on 22 Jun RELATED APPLN. INFO.:

1999

Utility DOCUMENT TYPE: GRANTED FILE SEGMENT:

Brusca, John S. PRIMARY EXAMINER: ASSISTANT EXAMINER: Lundgren, Jeffrey S.

Seed Intellectual Property Law Group PLLC LEGAL REPRESENTATIVE:

24 NUMBER OF CLAIMS: EXEMPLARY CLAIM:

APPLICATION INFO.:

NUMBER OF DRAWINGS: 17 Drawing Figure(s); 11 Drawing Page(s)

LINE COUNT: 4803

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2001 ACS

Method of affinity purifying proteins using modified bis-arsenical TI

The present invention features methods for purifying polypeptides of ABinterest using a modified Fluorescein arsenical helix binder (FlAsH) compd. immobilized on a solid support. An exemplary FlAsH target sequence motif is also presented. Examples of modification of the FlAsH compd. which allow immobilization

a solid support are also provided. The present invention also provides DNA constructs for producing a dual affinity tagged polypeptide and methods for purifn. thereof. Human kinesin constructs C-terminally tagged

with the peptide WEAAAREACCRECCARA (specifically chelating with .beta.-alanine-modified FlAsH, prepn. given) were expressed in Escherichia

coli and purified using beads contg. .beta.-alanine-modified FlAsH. Protein was eluted using 1,2-ethanedithiol.

2001:545718 HCAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 135:149588

Method of affinity purifying proteins using modified TITLE:

bis-arsenical fluorescein

INVENTOR(S):

Vale, Ronald D.; Thorn, Kurt; Cooke, Roger; Matuska,
Marija; Naber, Nariman

PATENT ASSIGNEE(S):
Regents of the University of ifornia, USA

SOURCE:

PCT Int. Appl., 52 pp.

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

W: AU, CA, JP

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,

PT, SE, TR

PRIORITY APPLN. INFO.: US 2000-178054 P 20000124

CODEN: PIXXD2

US 2000-502664 A 20000211

OTHER SOURCE(S): MARPAT 135:149588

L3 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2001 ACS

TI A FLASH of insight into cellular chemistry: genetically encoded labels for

protein visualization in vivo

AB A review with 24 refs. Genetically encoded fluorescent labels, such as green fluorescent protein, make it possible to visualize a protein's natural distribution and environment in living cells. A new approach to protein labeling in living cells has been devised in which a small, membrane-permeable ligand binds with high affinity and specificity to a short peptide motif that can be incorporated into the protein of interest;

the ligand becomes brightly fluorescent after binding to the peptide.

ACCESSION NUMBER: 1998:802921 HCAPLUS

DOCUMENT NUMBER: 130:150455

TITLE: A FLASH of insight into cellular chemistry:

genetically encoded labels for protein visualization

in vivo

AUTHOR(S): Leubke, Kevin J.

CORPORATE SOURCE: Center for Biomedical Inventions, Department of

Internal Medicine, University of Texas Southwestern Medical Center at Dallas, Dallas, TX, 75235-8573, USA

SOURCE: Chem. Biol. (1998), 5(12), R317-R322

CODEN: CBOLE2; ISSN: 1074-5521

PUBLISHER: Current Biology Publications

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

REFERENCE COUNT: 24

REFERENCE(S): (1) Chalfie, M; Science 1994, V263, P802 HCAPLUS

(2) Cody, C; Biochemistry 1993, V32, P1212 HCAPLUS

(3) Cubitt, A; Trends Biochem Sci 1995, V20, P448

HCAPLUS

(4) Golovina, V; Science 1997, V275, P1643 HCAPLUS

(5) Griffin, B; Science 1998, V281, P269 HCAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> e vale, r/au

E1	1	VALE WYLIE	WALKER/AU
E2	37	VALE WYLIE	WALKER JR/AU
E3	0>	VALE, R/AU	,
E4	19	VALEA A/AU	
E5	2	VALEA D/AU	
E6	1	VALEA E/AU	

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E7
                   VALEA F/AU
            17
E8
            28
                   VALEA EA/AU
E9
             4
                   VALEA
                          EL/AU
E10
             3
                   VALEA FIDEL A/AU
E11
                   VALEA GHEORGHE/AU
E12
                   VALEA GREGORIO/AU
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=> e cook, r/au
                   COOK YALE B/AU
E1
             1
E2
                   COOK YARWORTH J/AU
             1
E3
             0 --> COOK, R/AU
E4
                   COOKAND D R/AU
             1
E5
             1
                   COOKE/AU
E6
           446
                   COOKE A/AU
E7
                   COOKE A C/AU
             4
E8
             2
                   COOKE A D/AU
E9
             1
                   COOKE A D A/AU
E10
            7
                   COOKE A F/AU
E11
            65
                   COOKE A H/AU
E12
            12
                   COOKE A I/AU
=> s fluorescein arsenical helix binder () solid support
             O FLUORESCEIN ARSENICAL HELIX BINDER (W) SOLID SUPPORT
L9
```

CN 5: PN: WO0047220 SEQID: 49 unclaimed sequence

FS PROTEIN SEQUENCE; STEREOSEARCH

SQL 17

SEQ 1 AEAAAREACC RECCARA

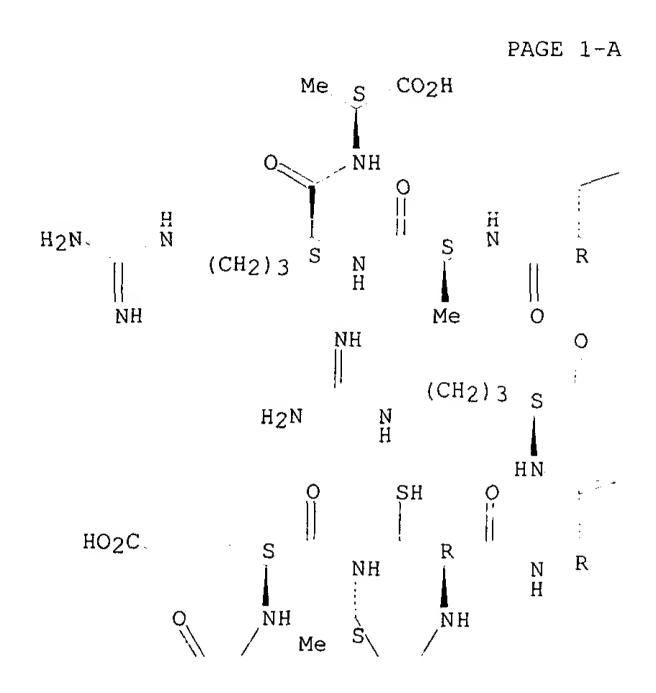
HITS AT: 9-14

MF C66 H114 N26 O24 S4

SR CA

LC STN Files: CA, CAPLUS, TOXLIT, USPATFULL

Absolute stereochemistry.



PAGE 1-B

2 REFERENCES IN FILE CA (1967 TO DATE) 2 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 133:172215

2: 130:308804 REFERENCE

L57 ANSWER 4 OF 4 REGISTRY COPYRIGHT 2001 ACS

RN**223673-78-7** REGISTRY

L-Alaninamide, N-acetyl-L-tryptophyl-L-.alpha.-glutamyl-L-alanyl-L-alanyl-CN L-alanyl-L-arginyl-L-.alpha.-glutamyl-L-alanyl-L-cysteinyl-L-cysteinyl-Larginyl-L-.alpha.-glutamyl-L-cysteinyl-L-cysteinyl-L-alanyl-L-arginyl-(9CI) (CA INDEX NAME)

PROTEIN SEQUENCE; STEREOSEARCH FS

SQL 17

modified NTE

type	locati	on	description	
terminal mod. terminal mod.	Trp-1 Ala-17	- -	N-acetyl C-terminal amide	

SEQ 1 WEAAAREACC RECCARA

9-14

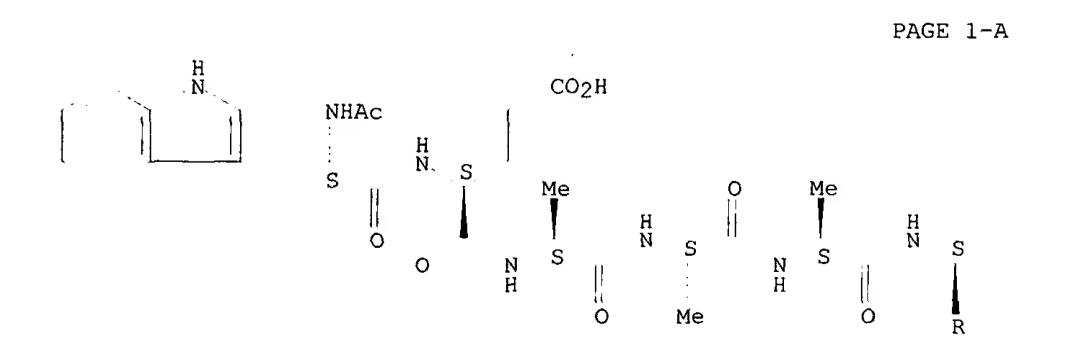
HITS AT:

C76 H122 N28 O24 S4 MF

SR CA

LCSTN Files: CA, CAPLUS, USPATFULL

Absolute stereochemistry.



PAGE 1-B

PAGE 2-B

1 REFERENCES IN FILE CA (1967 TO DATE)
1 REFERENCES IN FILE CAPLUS (1967 TO DATE)

REFERENCE 1: 130:308804

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FILE COVERS 1947 - 13 Nov 2001 VOL 135 ISS 21 FILE LAST UPDATED: 12 Nov 2001 (20011112/ED)
```

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```
=> d all tot 151
    ANSWER 1 OF 9 HCAPLUS
L51
                             COPYRIGHT 2001 ACS
     2001:545718 HCAPLUS
AN
DN
     135:149588
     Method of affinity purifying proteins using modified bis-arsenical
TI
     fluorescein
     Vale, Ronald D.; Thorn, Kurt; Cooke, Roger;
IN
     Matuska, Marija; Naber, Nariman
     The Regents of the University of California, USA
PA
SO
     PCT Int. Appl., 52 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
     ICM CO7K
IC
     9-3 (Biochemical Methods)
CC
     Section cross-reference(s): 28
FAN.CNT 1
     PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
     WO 2001053325
                       A2
PΙ
                            20010726
                                           WO 2001-US2214
                                                            20010122
         W: AU, CA, JP
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
PRAI US 2000-178054
                            20000124
     US 2000-502664
                            20000211
OS
     MARPAT 135:149588
     The present invention features methods for purifying polypeptides of
AB
     interest using a modified Fluorescein arsenical helix
     binder (FlAsH) compd. immobilized on a solid support. An exemplary FlAsH
     target sequence motif is also presented. Examples of modification of the
     FlAsH compd. which allow immobilization to a solid support are also
     provided. The present invention also provides DNA constructs for
     producing a dual affinity tagged polypeptide and methods for purifn.
     thereof. Human kinesin constructs C-terminally tagged with the peptide
     WEAAAREACCRECCARA (specifically chelating with .beta. -
     alanine-modified FlAsH, prepn. given) were expressed in
     Escherichia coli and purified using beads contg. .beta.-
     alanine-modified FlAsH. Protein was eluted using
     1,2-ethanedithiol.
ST
    protein purifn bis arsenical fluorescein helix binder;
     immobilization affinity purifn protein fluorescein
    arsenical compd; beta alanine modified bis
    arsenical fluorescein; kinesin fusion protein purifn
    arsenical fluorescein
IT
    Liquid chromatography
        (FPLC; affinity purifying proteins using modified bis-arsenical
       fluorescein)
    Affinity chromatographic stationary phases
ΙT
        (HPLC; affinity purifying proteins using modified bis-arsenical
```

```
fluorescein)
     Proteins, specific or class
IT
     RL: BPN (Biosynthetic preparation); NUU (Nonbiological use, unclassified);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
         (MBP (maltose-binding protein), chimeric protein contq.; affinity
        purifying proteins using modified bis-arsenical
        fluorescein)
     Affinity chromatographic stationary phases
IT
     Affinity chromatography
     Gel permeation chromatography
     Immobilization, biochemical
     Molecular cloning
     Plant cell
     рΗ
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
     Peptides, biological studies
ΙT
     Proteins, general, biological studies
     RL: BPN (Biosynthetic preparation); BPR (Biological process); PUR
     (Purification or recovery); BIOL (Biological study); PREP (Preparation);
     PROC (Process)
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
     Fusion proteins (chimeric proteins)
ΙT
     RL: BPN (Biosynthetic preparation); NUU (Nonbiological use, unclassified);
     PUR (Purification or recovery); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
IΤ
     Kinesins
     RL: BPN (Biosynthetic preparation); PUR (Purification or recovery); BIOL
     (Biological study); PREP (Preparation)
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
IT
     HPLC
     HPLC stationary phases
     Purification
        (affinity; affinity purifying proteins using modified bis-
        arsenical fluorescein)
     Ceramics
     Latex
     Paper
        (as support; affinity purifying proteins using modified bis-
        arsenical fluorescein)
     Fluoropolymers, uses
IT
    Metals, uses
     Plastics, uses
     Polyamides, uses
     Polyesters, uses
     Polymers, uses
     Polysiloxanes, uses
     Rayon, uses
     Semimetals
    RL: DEV (Device component use); USES (Uses)
        (as support; affinity purifying proteins using modified bis-
        arsenical fluorescein)
ΙT
    Chromatography
        (batch; affinity purifying proteins using modified bis-
        arsenical fluorescein)
    Insect (Insecta)
IT
        (cells of; affinity purifying proteins using modified bis-
        arsenical fluorescein)
    Polymers, uses
IT
    RL: DEV (Device component use); USES (Uses)
        (co-, as support; affinity purifying proteins using modified bis-
        arsenical fluorescein)
```

```
IT
     DNA
     RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP
     (Preparation)
        (constructs for producing dual affinity tagged polypeptides; affinity
        purifying proteins using modified bis-arsenical
        fluorescein)
    Glass, uses
IT
     RL: DEV (Device component use); USES (Uses)
        (controlled pore, as support; affinity purifying proteins using
        modified bis-arsenical fluorescein)
     Thiols (organic), uses
\operatorname{IT}
     RL: NUU (Nonbiological use, unclassified); USES (Uses)
        (dithiols, polypeptide elution with; affinity purifying proteins using
        modified bis-arsenical fluorescein)
     Proteins, specific or class
IT
     RL: BPN (Biosynthetic preparation); PUR (Purification or recovery); BIOL
     (Biological study); PREP (Preparation)
        (dual affinity-tagged; affinity purifying proteins using modified bis-
        arsenical fluorescein)
    Amino acids, reactions
IT
     RL: RCT (Reactant)
        (fluorescein arsenical helix binder modification by
        acylation with; affinity purifying proteins using modified bis-
        arsenical fluorescein)
IT
    Acylation
        (fluorescein arsenical helix binder modification by
        amino acid; affinity purifying proteins using modified bis-
        arsenical fluorescein)
     Proteins, specific or class
ΙT
     RL: BPN (Biosynthetic preparation); BUU (Biological use, unclassified);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (fluorescent, gene for, as selectable marker in DNA construct; affinity
        purifying proteins using modified bis-arsenical
       fluorescein)
    Chimeric gene
ΙT
     RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP
     (Preparation)
        (for affinity tag and fluorescein arsenical helix
        binder target sequence motif; affinity purifying proteins using
       modified bis-arsenical fluorescein)
     Protein motifs
IT
        (for fluorescein arsenical helix binder target
        sequence; affinity purifying proteins using modified bis-
       arsenical fluorescein)
IT
    Gene
     RL: BPN (Biosynthetic preparation); BUU (Biological use, unclassified);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (for selectable marker, in DNA construct; affinity purifying proteins
       using modified bis-arsenical fluorescein)
    Antibiotic resistance
IT
        (gene for, as selectable marker in DNA construct; affinity purifying
       proteins using modified bis-arsenical fluorescein)
IT
     Toxins
    RL: BPN (Biosynthetic preparation); BUU (Biological use, unclassified);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene for, as selectable marker in DNA construct; affinity purifying
       proteins using modified bis-arsenical fluorescein)
IT
     Genetic markers
        (in DNA construct; affinity purifying proteins using modified bis-
        arsenical fluorescein)
     Escherichia coli
IT
        (kinesin tagged with FlAsH peptide target prodn. in and purifn. from;
       affinity purifying proteins using modified bis-arsenical
        fluorescein)
    Cell
ΙT
```

(lysates, polypeptides of; affinity purifying proteins using modified

```
bis-arsenical fluorescein)
     Eukaryote (Eukaryotae)
IT
     Plant (Embryophyta)
     Prokaryote
        (polypeptides of; affinity purifying proteins using modified bis-
        arsenical fluorescein)
     26062-48-6DP, Polyhistidine, chimeric proteins
                                                      50812-37-8DP, Glutathione
IT
     S-transferase, chimeric proteins 64134-30-1DP, (L-His)6, chimeric
                98849-88-8DP, FLAG peptide, chimeric proteins
     proteins
     RL: BPN (Biosynthetic preparation); NUU (Nonbiological use, unclassified);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
IT
     2321-07-5D, Fluorescein, modified arsenical
     helix binder, immobilized 268741-25-9D, tautomers and anhydrides
     and salts and immobilized
     RL: NUU (Nonbiological use, unclassified); USES (Uses)
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
IT
     268741-25-9P
     RL: NUU (Nonbiological use, unclassified); SPN (Synthetic preparation);
     PREP (Preparation); USES (Uses)
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
     3326-34-9, 4-Amino fluorescein 7784-34-1,
IT
                           35737-10-1
     Arsenic trichloride
     RL: RCT (Reactant)
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
    73857-22-4P
                   268741-26-0P
                                  268741-27-1P
IT
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)
        (affinity purifying proteins using modified bis-arsenical
        fluorescein)
     268741-28-2P
IT
     RL: BPN (Biosynthetic preparation); BPR (Biological process); NUU
     (Nonbiological use, unclassified); PRP (Properties); BIOL (Biological
     study); PREP (Preparation); PROC (Process); USES (Uses)
        (amino acid sequence, as FlAsH peptide target; affinity purifying
       proteins using modified bis-arsenical fluorescein)
     352206-69-0
IT
     RL: BPR (Biological process); NUU (Nonbiological use, unclassified); PRP
     (Properties); BIOL (Biological study); PROC (Process); USES (Uses)
        (amino acid sequence, fluorescein arsenical helix
        binder target sequence; affinity purifying proteins using modified bis-
        arsenical fluorescein)
     9002-81-7, Polyformaldehyde
                                   9002-84-0, Teflon
                                                       9002-88-4, Polyethylene
IT
     9003-07-0, Polypropylene
                              9003-53-6, Polystyrene
                                                         9004-34-6, Cellulose,
                                           9004-70-0, Nitrocellulose
            9004-35-7, Cellulose acetate
                         24937-79-9, Polyvinylidene difluoride 24991-31-9,
     9012-36-6, Agarose
                         25038-59-9, Poly(ethylene terephthalate), uses
     Polv(vinvlbutvrate)
     25087-26-7, Polymethacrylic acid 28902-82-1
                                                     34540-03-9, Polyacrylimide
     RL: DEV (Device component use); USES (Uses)
        (as support; affinity purifying proteins using modified bis-
        arsenical fluorescein)
\mathbf{IT}
    107-95-9, .beta.-Alanine
    RL: RCT (Reactant)
        (fluorescein arsenical helix binder modification by
        acylation with; affinity purifying proteins using modified bis-
        arsenical fluorescein)
              288-32-4, Imidazole, uses 3483-12-3,
IT
     74-61-3
     Dithiothreitol
     RL: NUU (Nonbiological use, unclassified); USES (Uses)
        (polypeptide elution with; affinity purifying proteins using modified
        bis-arsenical fluorescein)
    540-63-6, 1,2-Ethanedithiol
IT
     RL: NUU (Nonbiological use, unclassified); RCT (Reactant); USES (Uses)
```

(polypeptide elution with; affinity purifying proteins using modified bis-arsenical fluorescein)

```
ANSWER 2 OF 9 HCAPLUS COPYRIGHT 2001 ACS
 L51
     2001:228897 HCAPLUS
 AN
     134:261272
 DN
     Cell membrane-impermeable arsenoxide compounds, their
 TI
     preparation, pharmaceutical compositions, and therapeutic and diagnostic
     use
     Hogg, Philip John; Donoghue, Neil
IN
     Unisearch Limited, Australia
PA
     PCT Int. Appl., 122 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LA
IC
     ICM C07F009-20
     ICS C07F009-78; C07F009-74
CC
     1-12 (Pharmacology)
     Section cross-reference(s): 29, 63
FAN.CNT 1
     PATENT NO.
                      KIND
                            DATE
                                            APPLICATION NO.
PI
     WO 2001021628
                            20010329
                       A1
                                            WO 2000-AU1143
                                                             20000920
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
             HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
             LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
             SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRAI AU 1999-2967
                            19990920
                       Α
OS
     MARPAT 134:261272
     The invention discloses compds. A(LY)p, (A = .gtoreq.1 substantially
AB
     cell-membrane impermeable pendant group; L = linker and/or spacer; Y =
     .gtoreq.1 arsenoxide or arsenoxide equiv.; p = 1-10;
     sum total of C atoms in A and L together >6). Prepn. of e.g.
     4-[N-(S-glutathionylacetyl)amino]phenylarsenoxide is described,
     as are e.g. the antitumor activity, tumor imaging ability, and activity
     inhibiting HIV infection of compds. of the invention. Pharmaceutical
     formulations are also described.
     membrane impermeable arsenoxide compd prepn therapeutic;
ST
     diagnostic membrane impermeable arsenoxide compd prep; antitumor
     tumor imaging arsenoxide compd; HIV infection arsenoxide
     compd
     Fluorescent substances
IT
        (arsenoxide derivs.; substantially cell membrane-impermeable
        compd. and use thereof)
IT
     Amines, biological studies
     Amino acids, biological studies
     Oligosaccharides, biological studies
     Peptides, biological studies
     Proteins, general, biological studies
     Radionuclides, biological studies
     Transition metals, biological studies
     RL: BAC (Biological activity or effector, except adverse); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (arsenoxide derivs.; substantially cell membrane-impermeable
        compd. and use thereof)
     Drug delivery systems
IT
        (capsules; substantially cell membrane-impermeable compd. and use
        thereof)
IT
    Lung, neoplasm
        (carcinoma, imaging; substantially cell membrane-impermeable compd. and
        use thereof)
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Cell proliferation
IT
        (disease; substantially cell membrane-impermeable compd. and use
        thereof)
     Blood vessel
ΙT
        (endothelium, cell; substantially cell membrane-impermeable compd. and
        use thereof)
     Antitumor agents
IT
        (fibrosarcoma; substantially cell membrane-impermeable compd. and use
        thereof)
     Drug delivery systems
IT
        (inhalants; substantially cell membrane-impermeable compd. and use
        thereof)
     Cell proliferation
IT
     Lung, neoplasm
     Pancreas, neoplasm
        (inhibitors; substantially cell membrane-impermeable compd. and use
        thereof)
     Drug delivery systems
IT
        (injections; substantially cell membrane-impermeable compd. and use
        thereof)
     Drug delivery systems
\mathbf{I}\mathbf{T}
        (lotions; substantially cell membrane-impermeable compd. and use
        thereof)
ΙŢ
     Antitumor agents
        (lung; substantially cell membrane-impermeable compd. and use thereof)
     Proteins, specific or class
IT
     RL: BAC (Biological activity or effector, except adverse); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (mercapto-contg., arsenoxide derivs.; substantially cell
        membrane-impermeable compd. and use thereof)
     Drug delivery systems
IT
        (ointments, creams; substantially cell membrane-impermeable compd. and
        use thereof)
     Drug delivery systems
IT
        (ointments; substantially cell membrane-impermeable compd. and use
        thereof)
    Antitumor agents
ΙT
        (pancreas; substantially cell membrane-impermeable compd. and use
        thereof)
    Drug delivery systems
        (parenterals; substantially cell membrane-impermeable compd. and use
        thereof)
     Drug delivery systems
IT
        (solns., ophthalmic; substantially cell membrane-impermeable compd. and
        use thereof)
IT
    Angiogenesis inhibitors
    Anti-AIDS agents
    Anti-inflammatory agents
    Anticoagulants
    Antitumor agents
    Antiviral agents
    Autoimmune disease
     Blood vessel, disease
    CD4-positive T cell
    Cardiovascular agents
     Cell membrane
     Drug delivery systems
     Human immunodeficiency virus 1
     Imaging agents
        (substantially cell membrane-impermeable compd. and use thereof)
IT
     Thioredoxins
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological
     process); PEP (Physical, engineering or chemical process); BIOL
     (Biological study); PROC (Process)
        (substantially cell membrane-impermeable compd. and use thereof)
    CD4 (antigen)
IT
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RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (substantially cell membrane-impermeable compd. and use thereof)
IT
     Drug delivery systems
        (topical; substantially cell membrane-impermeable compd. and use
        thereof)
     14133-76-7, Technetium-99, biological studies
IT
     RL: BAC (Biological activity or effector, except adverse); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (metastable, arsenoxide derivs.; substantially cell
        membrane-impermeable compd. and use thereof)
     1119-62-6P 1122-90-3P 51146-91-9P
IT
                                         57757-57-0P
                    331722-83-9P 331722-84-0P 331722-85-1P
     331722-76-0P
     331722-86-2P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)
        (prepn. and reaction; substantially cell membrane-impermeable compd.
        and use thereof)
     56-84-8, L-Aspartic acid, reactions 56-86-0, L-Glutamic acid, reactions
IT
     66-84-2, D-Glucosamine hydrochloride 70-18-8, Glutathione, reactions
     98-50-0, p-Arsanilic acid 107-96-0, 3-Mercaptopropanoic acid
     498-40-8, L-Cysteic acid 598-21-0, Bromoacetyl bromide
     N-Hydroxysuccinimide 67278-31-3 89889-52-1 123740-08-9
                  148356-01-8 172777-84-3, Cy5.5
     148356-00-7
     RL: RCT (Reactant)
        (reaction; substantially cell membrane-impermeable compd. and use
        thereof)
IT
     331722-70-4P
     RL: BAC (Biological activity or effector, except adverse); PEP (Physical,
     engineering or chemical process); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC
     (Process); USES (Uses)
        (substantially cell membrane-impermeable compd. and use thereof)
     331722-77-1P 331722-78-2P
IT
                                331722-79-3P
     331722-80-6P
                   331722-81-7P
                                  331722-82-8P 331722-87-3P
     331722-88-4P 331722-89-5P 331722-90-8P
    RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic
    preparation); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (substantially cell membrane-impermeable compd. and use thereof)
    56-84-8D, Aspartic acid, arsenoxide derivs. 56-86-0D, Glutamic
ΙT
    acid, arsenoxide derivs. 56-87-1D, Lysine, arsenoxide
              58-85-5D, Biotin, arsenoxide derivs.
    Glutathione, arsenoxide derivs. 74-79-3D, Arginine,
    arsenoxide derivs. 498-40-8D, Cysteic acid, arsenoxide
    derivs. 2321-07-5D, Fluorescein, arsenoxide
              3416-24-8D, Glucosamine, arsenoxide derivs.
    7440-38-2D, Arsenic, arsenoxide derivs.,
    biological studies 10028-17-8D, Tritium, arsenoxide derivs.,
    biological studies 10043-66-0D, Iodine-131, arsenoxide
    derivs., biological studies 13967-65-2D, Holmium-166, arsenoxide
    derivs., biological studies
                                  14119-09-6D, Gallium-67, arsenoxide
    derivs., biological studies 14158-31-7D, Iodine-125, arsenoxide
    derivs., biological studies 14265-75-9D, Lutetium-177,
    arsenoxide derivs., biological studies 14378-26-8D, Rhenium-188,
    arsenoxide derivs., biological studies 14596-37-3D,
    Phosphorus-32, arsenoxide derivs., biological studies
    14762-75-5D, Carbon-14, arsenoxide derivs., biological studies
    14913-89-4D, arsenoxide derivs., biological studies
    14998-63-1D, Rhenium-186, arsenoxide derivs., biological studies
    15117-53-0D, Sulfur-35, arsenoxide derivs., biological studies
    15715-08-9D, Iodine-123, arsenoxide derivs., biological studies
    15749-66-3D, Phosphorus-33, arsenoxide derivs., biological
    studies 15750-15-9D, Indium-111, arsenoxide derivs.,
    biological studies 15757-86-5D, Copper-67, arsenoxide derivs.,
    biological studies 15766-00-4D, Samarium-153, arsenoxide
    derivs., biological studies 19246-18-5D, Cysteinylglycine,
    arsenoxide derivs.
                        172777-84-3D, Cy5.5, arsenoxide
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derivs. 331722-71-5 331722-72-6 331722-73-7
     331722-74-8 331746-49-7 331815-00-0
     331815-01-1 331815-02-2 331815-03-3
     331815-04-4 331815-05-5 331815-06-6
     331815-07-7 331815-08-8 331815-09-9
     331815-10-2
     RL: BAC (Biological activity or effector, except adverse); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (substantially cell membrane-impermeable compd. and use thereof)
     37318-49-3, Protein disulfide isomerase
IT
     RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (substantially cell membrane-impermeable compd. and use thereof)
     59-52-9, 2,3-Dimercapto-1-propanol 69-78-3, DTNB
IT
     6,8-Thioctic acid 3483-12-3, Dithiothreitol
     RL: PEP (Physical, engineering or chemical process); PROC (Process)
        (substantially cell membrane-impermeable compd. and use thereof)
                   331722-91-9
IT
     117525-19-6
     RL: PEP (Physical, engineering or chemical process); PRP (Properties);
     PROC (Process)
        (substantially cell membrane-impermeable compd. and use thereof)
RE.CNT 7
RE
(1) Bhargava; Mol Biochem Parasitol 1983, V9, P29 HCAPLUS
(2) Carter; Nature 1993, V361(6408), P173 HCAPLUS
(3) Cunningham; Eur J Biochem 1994, V221, P285 HCAPLUS
(4) Fairlamb; Proc Natl Acad Sci 1989, V86, P2607 HCAPLUS
(5) Fairlamb, A; Ann Rev Microbiol 1992, V46, P695 HCAPLUS
(6) Friedheim; US 3883650 1975 HCAPLUS
(7) Pisciotto; Biochimica et Biophysica acta 1980, V628, P241 HCAPLUS
L51 ANSWER 3 OF 9 HCAPLUS COPYRIGHT 2001 ACS
AN
     2000:911534 HCAPLUS
DN
     134:66121
    Compositions and methods for assaying subcellular conditions and processes
TI
     using energy transfer for drug screening
     Dykens, James A.; Velicelebi, Gonul; Ghosh, Soumitra S.
IN
     Mitokor, USA
PΑ
     PCT Int. Appl., 189 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
ĽΑ
IC
     ICM G01N033-50
CC
     1-1 (Pharmacology)
FAN.CNT 1
     PATENT NO.
                                           APPLICATION NO.
                      KIND
                            DATE
                                                            DATE
                       A2
                            20001228
                                           WO 2000-US17380
     WO 2000079274
PI
                                                            20000622
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             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
             HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
             LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
             SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
             YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     US 6280981
                                           US 2000-514569
                       B1
                            20010828
                                                            20000223
PRAI US 1999-140433
                       Ρ
                            19990622
                            19990622
     US 1999-338122
                       Α
     US 2000-176383
                       Ρ
                            20000114
     The invention provides compns. and methods for monitoring subcellular
AB
     compartments such as organelles by energy transfer techniques that do not
     require specific intermol. affinity binding events between energy transfer
     donor and energy transfer acceptor mols. pH. Provided are methods for
     assaying cellular membrane potential, including mitochondrial membrane
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potential, by energy transfer methodologies including fluorescence

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resonance energy transfer (FRET). Diagnostic and drug screening assays
      are also provided.
      fluorescence resonance energy transfer FRET drug screening cell
 ST
      mitochondrium
      Transport proteins
 IT
      RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
         (ADP/ATP carrier; compns. and methods for assaying subcellular
         conditions and processes using energy transfer for drug screening)
 IT
      Fluorescent probes
         (LysoSensor and LysoTracker; compns. and methods for assaying
         subcellular conditions and processes using energy transfer for drug
        screening)
 IT
     Membrane potential
         (biol.; compns. and methods for assaying subcellular conditions and
        processes using energy transfer for drug screening)
     Alzheimer's disease
 IT
     Animal tissue culture
     Apoptosis
     Drug screening
     Fluorometry
     Ion channel blockers
     Mitochondria
     Parkinson's disease
     Permeability
     Plant tissue culture
     Hq
        (compns. and methods for assaying subcellular conditions and processes
        using energy transfer for drug screening)
     Natural products, pharmaceutical
ΙT
     RL: BAC (Biological activity or effector, except adverse); BIOL
     (Biological study)
        (compns. and methods for assaying subcellular conditions and processes
        using energy transfer for drug screening)
     Calcium channel
ΙT
     RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (compns. and methods for assaying subcellular conditions and processes
        using energy transfer for drug screening)
ΙT
     Glutamate receptors
     RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (compns. and methods for assaying subcellular conditions and processes
        using energy transfer for drug screening)
     Resonant energy transfer
IT
        (fluorescence; compns. and methods for assaying subcellular conditions
        and processes using energy transfer for drug screening)
IT
     Proteins, specific or class
     RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); ANST
     (Analytical study); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (green fluorescent, blue shifted; compns. and methods for assaying
        subcellular conditions and processes using energy transfer for drug
        screening)
     Proteins, specific or class
ΙT
     RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); ANST
     (Analytical study); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (green fluorescent, cyan shifted; compns. and methods for assaying
        subcellular conditions and processes using energy transfer for drug
        screening)
     Proteins, specific or class
IT
     RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);
    ANST (Analytical study); BIOL (Biological study); USES (Uses)
        (green fluorescent, red shifted; compns. and methods for assaying
        subcellular conditions and processes using energy transfer for drug
        screening)
     Proteins, specific or class
IT
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RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);

IT

TI

ΙT

TI

IT

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IT

IT

IT

IT

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ANST (Analytical study); BIOL (Biological study); USES (Uses)
   (green fluorescent, yellow shifted; compns. and methods for assaying
   subcellular conditions and processes using energy transfer for drug
   screening)
Proteins, specific or class
RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);
ANST (Analytical study); BIOL (Biological study); USES (Uses)
   (green fluorescent; compns. and methods for assaying subcellular
   conditions and processes using energy transfer for drug screening)
Polarization
   (hyperpolarization, biol., of mitochondria; compns. and methods for
   assaying subcellular conditions and processes using energy transfer for
   drug screening)
Mitochondria
   (membrane; compns. and methods for assaying subcellular conditions and
   processes using energy transfer for drug screening)
Membrane, biological
   (mitochondrial; compns. and methods for assaying subcellular conditions
   and processes using energy transfer for drug screening)
Diabetes mellitus
   (non-insulin-dependent; compns. and methods for assaying subcellular
   conditions and processes using energy transfer for drug screening)
199116-50-2, MitoTracker Orange CMTMRos
RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);
ANST (Analytical study); BIOL (Biological study); USES (Uses)
   (MitoTracker Orange CMTMRos; compns. and methods for assaying
   subcellular conditions and processes using energy transfer for drug
   screening)
                                 989-38-8, Rhodamine 6G 1239-45-8,
81-88-9, Rhodamine B 959-81-9
Ethidium bromide 2156-29-8 2315-97-1, Lucigenin 3520-43-2, JC-1
3785-01-1, DASPEI 6837-70-3, Rosamine 14806-50-9 41085-99-8
47165-04-8, DAPI 53213-81-3 53213-82-4 53213-83-5
                                                         59865-13-3,
Cyclosporin A 62669-70-9, Rhodamine 123 75168-11-5, 10-Nonyl acridine
       84109-11-5 86701-10-2 94885-04-8 115532-49-5,
orange
Tetramethylrhodamine, methyl ester 139626-15-6, Tetramethylrhodamine
ethylester 161057-69-8, FUN-1 201860-17-5, MitoTracker Green FM
212118-77-9, Spiro[isobenzofuran-1(3H),9'-[9H]xanthen]-3-one,
4',5'-bis(1,3,2-dithiarsolan-2-yl)-3',6'-dihydroxy- 273720-46-0,
                 314266-84-7, SNAFL calcein 314266-85-8 314730-55-7,
MitoFluor green
SYTO 18
RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);
ANST (Analytical study); BIOL (Biological study); USES (Uses)
   (compns. and methods for assaying subcellular conditions and processes
   using energy transfer for drug screening)
56-86-0, L-Glutamic acid, biological studies 370-86-5, Carbonyl cyanide
p-(trifluoromethoxy)phenyl hydrazone 487-79-6, Kainic acid 555-60-2,
Carbonyl cyanide m-chlorophenyl hydrazone 1404-19-9, Oligomycin
3106-85-2, NAAG 6384-92-5, NMDA 11076-19-0, Bongkrekic acid
17754-44-8, Atractyloside 28380-24-7, Nigericin
                                                   33286-30-5,
Carboxyatractyloside 48134-75-4, 1-Methyl-4-phenylpyridinium
52665-69-7, A23187 60132-21-0, Isobongkrekic acid 67526-95-8,
Thapsigargin 77521-29-0, 4-Isoxazolepropanoic acid, .alpha.-amino-2,3-
dihydro-5-methyl-3-oxo- 154461-69-5
RL: BAC (Biological activity or effector, except adverse); BIOL
(Biological study)
   (compns. and methods for assaying subcellular conditions and processes
   using energy transfer for drug screening)
7440-70-2, Calcium, biological studies
RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
   (compns. and methods for assaying subcellular conditions and processes
   using energy transfer for drug screening)
25125-46-6
RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);
ANST (Analytical study); BIOL (Biological study); USES (Uses)
   (ruthenium red; compns. and methods for assaying subcellular conditions
   and processes using energy transfer for drug screening)
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83796-96-7, Tetrabromo-rhodamine 123
 IT
      RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);
      ANST (Analytical study); BIOL (Biological study); USES (Uses)
         (tetrabromorhodamine 123; compns. and methods for assaying subcellular
         conditions and processes using energy transfer for drug screening)
     ANSWER 4 OF 9 HCAPLUS COPYRIGHT 2001 ACS
 L51
     2000:861509 HCAPLUS
 AN
     134:27295
 DN
     Methods for producing 5'-nucleic acid-protein conjugates
 TI
     Lohse, Peter; Wright, Martin C.; McPherson, Michael
 IN
 PA
     Phylos, Inc., USA
     PCT Int. Appl., 32 pp.
 SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
     ICM A61K038-16
IC
     ICS A61K038-03; C07K014-00
     9-16 (Biochemical Methods)
CC
     Section cross-reference(s): 33, 34
FAN.CNT 1
     PATENT NO.
                      KIND
                            DATE
                                           APPLICATION NO.
                            20001207
PI
     WO 2000072869
                     A1
                                           WO 2000-US15077 20000601
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
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             ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
             LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
             SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW,
             AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRAI US 1999-137032
                           19990601
                      P
     Disclosed herein is a method for generating a 5'-nucleic acid-protein
AΒ
     conjugate, the method involving: (a) providing a nucleic acid which
     carries a reactive group at its 5'end; (b) providing a non-derivatized
     protein; and (c) contacting the nucleic acid and the protein under
     conditions which allow the reactive group to react with the N-terminus of
     the protein, thereby forming a 5'-nucleic acid-protein conjugate. In one
     approach, fusions are formed by reaction between an unprotected protein
     carrying an N-terminal cysteine and a nucleic acid carrying a
     1,2-aminothiol reactive group. In a second approach, fusion formation
     occurs as the result of a biarsenical-tetracysteine interaction. Also
     disclosed herein are 5'-nucleic acid-protein conjugates and methods for
     their use in (1) the selection of a desired nucleic acid or a desired
     protein by sepg. the binding partner-candidate conjugate complex from
     unbound members of a population, and (2) detecting an interaction between
     a protein and a compd.
     nucleic acid conjugate protein
ST
IT
     Thiols (organic), reactions
     RL: RCT (Reactant)
        (amino, nucleic acids contg., reaction with N-terminal cysteinyl
       proteins; methods for producing 5'-nucleic acid-protein conjugates)
IT
     DNA
    Nucleic acids
     Proteins, specific or class
     RNA
    mRNA
    RL: ANT (Analyte); BUU (Biological use, unclassified); SPN (Synthetic
    preparation); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (conjugates; methods for producing 5'-nucleic acid-protein conjugates)
    Nucleoproteins
IT
    RL: BUU (Biological use, unclassified); SPN (Synthetic preparation); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
```

(methods for producing 5'-nucleic acid-protein conjugates) Amines, reactions IT RL: RCT (Reactant) (thiol, nucleic acids contg., reaction with N-terminal cysteinyl proteins; methods for producing 5'-nucleic acid-protein conjugates) 56377-57-2 ITRL: RCT (Reactant) (methods for producing 5'-nucleic acid-protein conjugates) 311797-38-3P 311797-39-4P IT RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (methods for producing 5'-nucleic acid-protein conjugates) 52-90-4, L-Cysteine, reactions IΤ RL: RCT (Reactant) (reaction with nucleic acid carrying a 1,2-aminothiol reactive group; methods for producing 5'-nucleic acid-protein conjugates) 312323-65-2 312323-67-4 312323-66-3 312343-85-4 ΙT RL: PRP (Properties) (unclaimed sequence; methods for producing 5'-nucleic acid-protein conjugates) RE.CNT 1 RE (1) Lebleu; EP 0263740 A1 1988 HCAPLUS L51 ANSWER 5 OF 9 HCAPLUS COPYRIGHT 2001 ACS AN 2000:850091 HCAPLUS 134:175077 DN Fluorescent labeling of recombinant proteins in living cells with FlAsH TIGriffin, B. Albert; Adams, Stephen R.; Jones, Jay; Tsien, Roger Y. ΑU Aurora Biosciences Corporation, San Diego, CA, 92121, USA CS Methods Enzymol. (2000), 327 (Applications of Chimeric Genes and Hybrid SO Proteins, Pt. B), 565-578 CODEN: MENZAU; ISSN: 0076-6879 Academic Press PBJournal DTEnglish LACC 9-4 (Biochemical Methods) Section cross-reference(s): 6 Chem. labeling of specific sites in proteins is usually achieved by ABreaction of single cysteine residues with appropriate thiol-reactive derivs. One approach for site-specific labeling of proteins in living cells is to utilize the well-known affinity of arsenoxides for a pair of closely spaced cysteines. To prevent labeling of such endogenous cellular sites, a fluorescein contg. two arsenoxides (FlAsH) was designed that has a much higher affinity for four appropriately spaced cysteines (CCXXCC, where X is any amino acid other than cysteine) in an .alpha.-helical conformation. Such motifs are sufficiently uncommon in naturally occurring proteins to permit specific modification of the target protein incorporating the introduced FlAsH site in living cells. By labeling in the presence of the arsenoxide antidote 1,2-ethanediol (EDT), nonspecific labeling and toxicity can be minimized because EDT forms more stable complexes with arsenic than do pairs of cysteines. Moreover, FlAsH complexed with two EDT mols., is membrane permeable and nonfluorescent yet becomes brightly fluorescent on binding the CCXXCC site, thereby decreasing background signal from unbound dye during labeling. The tetracysteine site can be attached as an N- or C-terminal tag or incorporated into a known .alpha.-helical structure. Addn. of a high concn. of EDT reverses the binding of FlAsH to the tetracysteines, permitting reversible labeling. Chem. modification of the fluorescein moiety allows incorporation of different photochem. properties or use as a handle to target other small mols. to proteins modified with the FlAsH site. FlAsH may also be used to label purified proteins in vitro as an alternative to fluorescein iodoacetamide or maleimide reagents, with the advantage that the tetracysteine-binding site can be labeled without affecting single cysteines in the mol. (c) 2000 Academic Press.

FlAsH fluorescent label protein fluorescence

ST IT

Cell



robinson - 09 / 502664 Fluorescence Fluorescent indicators (fluorescent labeling of recombinant proteins in living cells with FlAsH) Proteins, general, analysis RL: ANT (Analyte); BSU (Biological study, unclassified); RCT (Reactant); ANST (Analytical study); BIOL (Biological study) (fluorescent labeling of recombinant proteins in living cells with FlAsH) 212118-77-9, FlAsH-EDT2 RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (FlAsH-EDT2; fluorescent labeling of recombinant proteins in living cells with FlAsH) 107-21-1, 1,2-Ethanediol, analysis RL: ARU (Analytical role, unclassified); ANST (Analytical study) (fluorescent labeling of recombinant proteins in living cells with FlAsH) 52-90-4, L-Cysteine, biological studies RL: BOC (Biological occurrence); RCT (Reactant); BIOL (Biological study); OCCU (Occurrence) (fluorescent labeling of recombinant proteins in living cells with FlAsH) RE.CNT 6 (1) Bachmair, A; Science 1986, V234, P179 HCAPLUS (2) Baker, R; J Biol Chem 1992, V267, P23364 HCAPLUS (3) Gilchrist, C; J Biol Chem 1997, V272, P32280 HCAPLUS (4) Tobias, J; J Biol Chem 1991, V266, P12021 HCAPLUS (5) Varshavsky, A; Proc Natl Acad Sci USA 1996, V93, P12142 HCAPLUS (6) Wilkinson, K; Ubiquitin and the Biology of the Cell 1998 ANSWER 6 OF 9 HCAPLUS COPYRIGHT 2001 ACS L51 2000:169797 HCAPLUS 132:344976 A novel method of affinity-purifying proteins using a bisarsenical fluorescein Thorn, Kurt S.; Naber, Nariman; Matuska, Marija; Vale, Ronald D.; Cooke, Roger Department of Cellular and Molecular Pharmacology, University of California, San Francisco, CA, 94143, USA Protein Sci. (2000), 9(2), 213-217 CODEN: PRCIEI; ISSN: 0961-8368 Cambridge University Press Journal English 9-3 (Biochemical Methods) Section cross-reference(s): 6 Genetically-encoded affinity tags constitute an important strategy for purifying proteins. Here, we have designed a novel affinity matrix based on the bis-arsenical fluorescein dye FlAsH, which specifically to FlAsH resin and can be eluted in a fully active form.

AΒ specifically recognizes short .alpha.-helical peptides contg. the sequence CCXXCC. We find that kinesin tagged with this cysteine-contg. helix binds This affinity tag has several advantages over polyhistidine, the only small affinity tag in common use. The protein obtained with this single chromatog, step from crude Escherichia coli lysates is purer than that obtained with nickel affinity chromatog. of 6xHis tagged kinesin. Moreover, unlike nickel affinity chromatog., which requires high concns. of imidazole or pH changes for elution, protein bound to the FlAsH column can be completely eluted by dithiothreitol. Because of these mild elution conditions, FlAsH affinity chromatog. is ideal for recovering fully active protein and for the purifn. of intact protein complexes.

protein kinesin affinity chromatog arsenical fluorescein ST

ΙT Affinity

ΙT

IT

IT

IT

RE

AN

DN

TI

ΑU

CS

SO

PBDT

LA

CC

Affinity chromatographic stationary phases Affinity chromatography

```
(novel method of affinity-purifying proteins using a bis-
        arsenical fluorescein)
IT
     Kinesins
     Proteins, general, biological studies
     RL: BSU (Biological study, unclassified); PUR (Purification or recovery);
     BIOL (Biological study); PREP (Preparation)
        (novel method of affinity-purifying proteins using a bis-
        arsenical fluorescein)
     268741-25-9P
IT
     RL: ARU (Analytical role, unclassified); BPR (Biological process); BUU
     (Biological use, unclassified); SPN (Synthetic preparation); ANST
     (Analytical study); BIOL (Biological study); PREP (Preparation); PROC
     (Process); USES (Uses)
        (novel method of affinity-purifying proteins using a bis-
        arsenical fluorescein)
     268741-28-2
IT
     RL: BPR (Biological process); BUU (Biological use, unclassified); BIOL
     (Biological study); PROC (Process); USES (Uses)
        (novel method of affinity-purifying proteins using a bis-
        arsenical fluorescein)
     3326-34-9, 4-Amino fluorescein 35737-10-1
IT
     RL: RCT (Reactant)
        (novel method of affinity-purifying proteins using a bis-
        arsenical fluorescein)
                                  268741-27-1P
     73857-22-4P
                   268741-26-0P
IT
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)
        (novel method of affinity-purifying proteins using a bis-
        arsenical fluorescein)
RE.CNT 10
RE
(1) Case, R; Cell 1997, V90, P959 HCAPLUS
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(3) Hannig, G; Trends Biotechnol 1998, V16, P54 HCAPLUS
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    ANSWER 7 OF 9 HCAPLUS COPYRIGHT 2001 ACS
L51
     1999:286159 HCAPLUS
NA
DN
     130:308804
     Target protein sequences for binding of synthetic biarsenical
TI
     molecules
     Tsien, Roger Y.; Griffin, Albert B.
ΙN
     The Regents of the University of California, USA
PA
     PCT Int. Appl., 77 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM G01N033-566
     ICS C07F009-80; C12N015-09; C12N015-64
CC
     9-15 (Biochemical Methods)
     Section cross-reference(s): 6
FAN. CNT 1
                                           APPLICATION NO.
                                                             DATE
     PATENT NO.
                      KIND
                            DATE
     WO 9921013
                                           WO 1998-US22363 19981021
                            19990429
                       A1
PΙ
        W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG,
             KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
             UA, UG, US, US, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
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RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
             FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
             CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                            19990803
     US 5932474
                       Α
                                           US 1997-955206
                                                            19971021
     US 6008378
                           19991228
                      Α
                                                            19971021
                                           US 1997-955859
                                           US 1997-955050
     US 6054271
                      A
                            20000425
                                                            19971021
                            19990510
    AU 9911139
                      A1
                                          AU 1999-11139
                                                            19981021
                            20000906
     EP 1032837
                                           EP 1998-953881
                       A1
                                                            19981021
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
PRAI US 1997-955050
                      A2
                            19971021
                      A2
     US 1997-955206
                           19971021
                      A2 19971021
     US 1997-955859
                            19981021
    WO 1998-US22363
                       W
    MARPAT 130:308804
os
    The present invention features biarsenical mols. and target
AB
     sequences that specifically react with the biarsenical mols. A
    bonding partner comprises a carrier polypeptide and a target sequence,
    wherein the target sequence is heterologous to the carrier polypeptide and
    the target sequence contains one or more cysteines capable of specifically
    reacting with a biarsenical mol. Bonding partners that include
    target sequences, vectors that include nucleic acid sequences that encode
    the target sequences and host cells that include the target sequences are
    also featured in the invention. One example of a biarsenical
    compd. is an arsenical deriv. of fluorescein.
    protein labeling crosslinking biarsenical mol
ST
IT
    Crosslinking agents
        (arsenic derivs.; target protein sequences for binding of
        synthetic biarsenical mols.)
    Proteins (specific proteins and subclasses)
IT
    RL: ANT (Analyte); BPR (Biological process); ANST (Analytical study); BIOL
     (Biological study); PROC (Process)
        (cyan fluorescent protein; target protein sequences for binding of
        synthetic biarsenical mols.)
    Peptides, analysis
IT
    Proteins (specific proteins and subclasses)
    RL: ARU (Analytical role, unclassified); BPR (Biological process); BUU
     (Biological use, unclassified); ANST (Analytical study); BIOL (Biological
    study); PROC (Process); USES (Uses)
        (labeled; target protein sequences for binding of synthetic
       biarsenical mols.)
    Antibodies
IT
    Enzymes, biological studies
    RL: BPR (Biological process); BUU (Biological use, unclassified); BIOL
     (Biological study); PROC (Process); USES (Uses)
        (labeled; target protein sequences for binding of synthetic
       biarsenical mols.)
    Green fluorescent protein
IT
    RL: ANT (Analyte); BPR (Biological process); ANST (Analytical study); BIOL
     (Biological study); PROC (Process)
        (mutant; target protein sequences for binding of synthetic
       biarsenical mols.)
IT
    Genes
    RL: BPR (Biological process); BUU (Biological use, unclassified); BIOL
     (Biological study); PROC (Process); USES (Uses)
        (recombinant; target protein sequences for binding of synthetic
       biarsenical mols.)
    Bacteria (Eubacteria)
ΙŢ
    Cell (biological)
    Conformation (protein)
    Crosslinking
    Fluorescence
    Fluorescent substances
    Genetic vectors
    HeLa cell
    Insect (Insecta)
```

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Labels
     Mammal (Mammalia)
                                                             (305, 505T)
     Mammalian cells
     Membranes (biological)
     Molecular association
     Plant (Embryophyta)
     Plant cells
     Protein sequences
     Yeast
     .alpha.-Helix (protein conformation)
        (target protein sequences for binding of synthetic biarsenical
        mols.)
    Calmodulins
IT
     Peptides, analysis
     Proteins (general), analysis
     RL: ANT (Analyte); BPR (Biological process); ANST (Analytical study); BIOL
     (Biological study); PROC (Process)
        (target protein sequences for binding of synthetic biarsenical
        mols.)
IT
     DNA
     RL: BPR (Biological process); BUU (Biological use, unclassified); BIOL
     (Biological study); PROC (Process); USES (Uses)
        (target protein sequences for binding of synthetic biarsenical
       mols.)
     223673-78-7
IT
    RL: ARU (Analytical role, unclassified); BPR (Biological process); ANST
     (Analytical study); BIOL (Biological study); PROC (Process)
        (SEQ ID 1; target protein sequences for binding of synthetic
       biarsenical mols.)
    223673-79-8
ΙT
    RL: ARU (Analytical role, unclassified); BPR (Biological process); ANST
     (Analytical study); BIOL (Biological study); PROC (Process)
        (SEQ ID 4; target protein sequences for binding of synthetic
       biarsenical mols.)
    7440-38-2D, Arsenic, derivs. 212118-77-9D,
IT
    tautomers, anhydrides and salts
    RL: ARG (Analytical reagent use); BPR (Biological process); ANST
     (Analytical study); BIOL (Biological study); PROC (Process); USES (Uses)
        (target protein sequences for binding of synthetic biarsenical
       mols.)
    212118-77-9P
IT
    RL: ARG (Analytical reagent use); BPR (Biological process); SPN (Synthetic
    preparation); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); PROC (Process); USES (Uses)
        (target protein sequences for binding of synthetic biarsenical
       mols.)
    52-90-4, L-Cysteine, biological studies
\operatorname{IT}
    RL: BOC (Biological occurrence); BPR (Biological process); PRP
     (Properties); BIOL (Biological study); OCCU (Occurrence); PROC (Process)
        (target protein sequences for binding of synthetic biarsenical
       mols.)
    223673-80-1 223673-81-2 223673-82-3
IT
    RL: BUU (Biological use, unclassified); RCT (Reactant); BIOL (Biological
    study); USES (Uses)
        (target protein sequences for binding of synthetic biarsenical
       mols.)
    76-54-0, 2',7'-Dichlorofluorescein 89-05-4, 1,2,4,5-
IT
    Benzenetetracarboxylic acid
                                   108-46-3, 1,3-Benzenediol, reactions
                                   1600-27-7, Mercuric acetate
    540-63-6, 1,2-Ethanedithiol
                                      32382-27-7,
    7784-34-1, Arsenic trichloride
                                    223673-84-5
    Fluorescein mercuric acetate
    RL: RCT (Reactant)
        (target protein sequences for binding of synthetic biarsenical
       mols.)
    54210-30-9P
                   223673-86-7P 223673-87-8P
IT
    RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation)
```

```
(target protein sequences for binding of synthetic biarsenical
        mols.)
TT
     223673-83-4P
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (target protein sequences for binding of synthetic biarsenical
        mols.)
RE.CNT
        2
RE
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L51 ANSWER 8 OF 9 HCAPLUS COPYRIGHT 2001 ACS
     1998:802921 HCAPLUS
AN
    130:150455
DN
TI
     A FLASH of insight into cellular chemistry: genetically encoded labels for
     protein visualization in vivo
     Leubke, Kevin J.
ΑŲ
     Center for Biomedical Inventions, Department of Internal Medicine,
CS
     University of Texas Southwestern Medical Center at Dallas, Dallas, TX,
     75235-8573, USA
     Chem. Biol. (1998), 5(12), R317-R322
SO
     CODEN: CBOLE2; ISSN: 1074-5521
     Current Biology Publications
PB
     Journal; General Review
\mathtt{T}\mathtt{T}
     English
LA
     9-0 (Biochemical Methods)
CC
     A review with 24 refs. Genetically encoded fluorescent labels, such as
AB
     green fluorescent protein, make it possible to visualize a protein's
     natural distribution and environment in living cells. A new approach to
     protein labeling in living cells has been devised in which a small,
     membrane-permeable ligand binds with high affinity and specificity to a
     short peptide motif that can be incorporated into the protein of interest;
     the ligand becomes brightly fluorescent after binding to the peptide.
     FLASH cell chem review; genetically encoded label protein review;
ST
     fluorescein arsenical helix binder review
     Fluorometry
ΙT
        (a FLASH of insight into cellular chem.: genetically encoded labels for
        protein visualization in vivo)
     Proteins (general), analysis
IT
     RL: ANT (Analyte); ANST (Analytical study)
        (a FLASH of insight into cellular chem.: genetically encoded labels for
        protein visualization in vivo)
     Green fluorescent protein
ΙT
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (a FLASH of insight into cellular chem.: genetically encoded labels for
        protein visualization in vivo)
IT
    Cell (biological)
        (living; a FLASH of insight into cellular chem.: genetically encoded
        labels for protein visualization in vivo)
     212118-77-9, Fluorescein arsenical helix binder
IT
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (FLASH; a FLASH of insight into cellular chem.: genetically encoded
        labels for protein visualization in vivo)
RE.CNT 24
RE
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- (23) Tsien, R; Ann Rev Neurosci 1989, V12, P227 HCAPLUS
- (24) Wang, S; Nature 1994, V369, P400 HCAPLUS
- L51 ANSWER 9 OF 9 HCAPLUS COPYRIGHT 2001 ACS
- AN 1998:456311 HCAPLUS
- DN 129:200138
- TI Specific covalent labeling of recombinant protein molecules inside live cells
- AU Griffin, B. Albert; Adams, Stephen R.; Tsein, Roger Y.
- CS Dep. Chem. and Biochem., Univ. California San diego, La Jolla, CA, 92093-0647, USA
- SO Science (Washington, D. C.) (1998), 281(5374), 269-272 CODEN: SCIEAS; ISSN: 0036-8075
- PB American Association for the Advancement of Science
- DT Journal
- LA English
- CC 9-16 (Biochemical Methods)
- AB Recombinant proteins contg. four cysteines at the i, i + 1, i + 4, and i + 5 positions of a .alpha. helix were fluorescently labeled in living cells by extracellular administration of 4',5'-bis(1,3,2-dithioarsolan-2-yl)fluorescein. This designed small ligand is membrane-permeant and nonfluorescent until it binds with high affinity and specificity to the tetracysteine domain. Such in situ labeling adds much less mass than does green fluorescent protein and offers greater versatility in attachment sites as well as potential spectroscopic and chem. properties. This system provides a recipe for slightly modifying a target protein so that it can be singled out from the many other proteins inside live cells and fluorescently stained by small nonfluorescent dye mols. added from outside the cells.
- ST covalent labeling recombinant protein live cell
- IT Cell (biological)
 - (live; specific covalent labeling of recombinant protein mols. inside live cells)
- IT Proteins (general), reactions
 - RL: RCT (Reactant)
 - (recombinant; specific covalent labeling of recombinant protein mols. inside live cells)
- IT .alpha.-Helix (protein conformation)
 - (specific covalent labeling of recombinant protein mols. inside live cells)
- IT 212118-77-9
 - RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (specific covalent labeling of recombinant protein mols. inside live cells)
- IT 52-90-4, Cysteine, reactions
 - RL: RCT (Reactant)
 - (specific covalent labeling of recombinant protein mols. inside live cells)

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FILE 'BIOSIS' ENTERED AT 16:04:07 ON 13 NOV 2001 COPYRIGHT (C) 2001 BIOSIS(R)

FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 7 November 2001 (20011107/ED)

The BIOSIS file has been reloaded. Enter HELP RLOAD and HELP REINDEXING for details.

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ANSWER 1 OF 2 BIOSIS COPYRIGHT 2001 BIOSIS L79

2000:201792 BIOSIS AN

PREV200000201792 DN

A novel method of affinity-purifying proteins using a bis-TIarsenical fluorescein.

Thorn, Kurt S.; Naber, Nariman; Matuska, ΑU Marija; Vale, Ronald D.; Cooke, Roger (1)

(1) Department of Biochemistry, University of California, San Francisco, CS 513 Parnassus Avenue, San Francisco, CA, 94143 USA

Protein Science, (Feb., 2000) Vol. 9, No. 2, pp. 213-217.
ISSN: 0961-8368.
Article
English SO

DT

LA

English SL

Genetically-encoded affinity tags constitute an important strategy for ABpurifying proteins. Here, we have designed a novel affinity matrix based on the bis-arsenical fluorescein dye FlAsH, which specifically recognizes short alpha-helical peptides containing the sequence CCXXCC (Griffin BA, Adams SR, Tsien RY, 1998, Science 281:269-272). We find that kinesin tagged with this cysteine-containing helix binds specifically to FlAsH resin and can be eluted in a fully active form. This affinity tag has several advantages over polyhistidine, the only small affinity tag in common use. The protein obtained with this single chromatographic step from crude Escherichia coli lysates is purer than that obtained with nickel affinity chromatography of 6xHis tagged kinesin. Moreover, unlike nickel affinity chromatography, which requires high concentrations of imidazole or pH changes for elution, protein bound to the FlAsH column can be completely eluted by dithiothreitol. Because of these mild elution conditions, FlAsH affinity chromatography is ideal for recovering fully active protein and for the purification of intact protein complexes.

Biochemical Studies - Proteins, Peptides and Amino Acids *10064 CC Biochemical Studies - General *10060 Biophysics - General Biophysical Techniques *10504 Physiology and Biochemistry of Bacteria *31000

Major Concepts IT

Biochemistry and Molecular Biophysics; Methods and Techniques

Chemicals & Biochemicals TI

Escherichia coli lysates: assay, purification; FIAsH: bisarsenical fluorescein dye, reagent, resin; proteins: assay, purification

ITMethods & Equipment

> FIAsH affinity chromatography: liquid chromatography, purification method; MALDI-MS [matrix-assisted laser/desorption ionization-mass spectrometry]: analytical method, mass spectrometry: CB; densitometry: analytical method, photometry: CB; nickel affinity chromatography: liquid chromatography, purification method

- ANSWER 2 OF 2 BIOSIS COPYRIGHT 2001 BIOSIS L79
- 1999:43454 BIOSIS AN
- PREV199900043454 DN
- A FLASH of insight into cellular chemistry: Genetically encoded \mathtt{TI} labels for protein visualization in vivo.
- Luebke, Kevin J. (1) ΑU
- (1) Cent. Biomedical Inventions, Dep. Internal Med., Univ. Texas CS

```
Southwestern Med. Cent. Dallas, 5323 Harry Hines Blvd., Dallas, TX
     75235-8573 USA
    Chemistry & Biology (London), (Dec., 1998) Vol. 5, No. 12, pp. R317-R322.
SO
     ISSN: 1074-5521.
    Article
\mathtt{DT}
    English
LA
    Genetically encoded fluorescent labels, such as green fluorescent
AB
    protein, make it possible to visualize a protein's
     natural distribution and environment in living cells. A new approach to
    protein labeling in living cells has been devised in which a
     small, membrane permeable ligand binds with high affinity and specificity
     to a short peptide motif that can be incorporated into the
    protein of interest; the ligand becomes brightly fluorescent after
    binding to the peptide.
CC Cytology and Cytochemistry - General *02502
    Methods, Materials and Apparatus, General - Laboratory Methods *01004
    Genetics and Cytogenetics - General *03502
     Radiation - General *06502
     Comparative Biochemistry, General *10010
     Biochemical Methods - General *10050
     Biochemical Studies - General *10060
     Biochemical Studies - Nucleic Acids, Purines and Pyrimidines *10062
       Biochemical Studies - Proteins, Peptides and Amino Acids *10064
     Replication, Transcription, Translation *10300
     Biophysics - Molecular Properties and Macromolecules *10506
     Biophysics - Membrane Phenomena *10508
    Metabolism - Proteins, Peptides and Amino Acids *13012
     In Vitro Studies, Cellular and Subcellular *32600
    Bacteria - General Unspecified
                                      05000
BC
    Animalia - Unspecified
                              33000
    Major Concepts
IT
        Biochemistry and Molecular Biophysics; Cell Biology; Methods and
        Techniques
     Parts, Structures, & Systems of Organisms
IT
        cell membranes
     Chemicals & Biochemicals
IT
          fluorescein arsenical helix binder: applications;
        fluorescent proteins: analysis; genetically-encoded
        fluorescent labels: applications; green fluorescent protein:
        applications; ligands; peptides; proteins:
        analysis, characterization, visualization
    Methods & Equipment
ΙT
        fluorescence energy transfer: Analysis/Characterization Techniques: CB,
        analytical method; protein labeling: Detection/Labeling
        Techniques, labeling method
    Miscellaneous Descriptors
\mathbf{T}\mathbf{T}
        cellular biochemistry; gene expression
ORGN Super Taxa
        Animalia; Bacteria: Microorganisms
ORGN Organism Name
        animals (Animalia); bacteria (Bacteria)
ORGN Organism Superterms
        Animals; Bacteria; Eubacteria; Microorganisms
RN
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L1
              1 S E3
                SEL RID
          10236 S E1
L2
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10 S L2 AND AS/ELS
L3
                 E .BETA.-ALANINE/CN
               1 S E3
L4
            807 S L2 AND SQL/FA
L5
            800 S L5 AND PROTEIN/FS
L6
               7 S L5 NOT L6
L7
               0 S L2 AND 107-95-9/CRN
\Gamma8
                 E 1,2-ETHANEDITHIOL/CN
L9
               1 S E3
                 E DITHIOTHERITOL/CN
L10
               1 S E4
                 E C4H10O2S2/MF
L11
               5 S E3 AND 2 3 BUTANEDIOL
                 E DIMERCAPTOPROPANESULFONATE/CN
                 E DIMERCAPTO PROPANESULFONATE/CN
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                 E THRON K/AU
                 E THORN K/AU
L13
             29 S E3-E5, E11, E12
                 E COOKE R/AU
            306 S E3-E20
L14
                 E COOKE ROGER/AU
            116 S E3-E6
L15
                 E MATUSKA M/AU
              4 S E3, E4
L16
                 E NABER N/AU
L17
             21 S E3-E6
L18
            570 S L12-L17
L19
           3815 S L1
          17627 S FLUORESCEIN
L20
L21
              4 S L18 AND L19, L20
L22
              2 S L21 AND ?ARSEN?
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L26
              7 S L24, L25
L27
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L28
              3 S L27 AND (6/SQL OR 17/SQL)
L29
              2 S L28 NOT NCNC2/ES
L30
              6 S L9-L11
              3 S L27 AND (C4H10O2S2 OR C3H8O3S3 OR C2H6S2)
L31
L32
              7 S L30, L31
L33
              4 S L3 AND 2/AS
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L38
L39
              2 S L37 AND L32
              2 S L37 AND L29
L40
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L41

837 S CC..CC/SQSP

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L42
              3 S L42 AND L37
L43
L44
              8 S L37-L40, L43
L45
             57 S L19, L20 AND ?ARSEN?
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L46
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             76 S L45, L47
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L50
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L51
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L53
              5 S L52 AND L1, L2
              1 S L52 AND L4
L54
L55
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                E THORN K/AU
             40 S E3-E5, E9, E10
L59
                E COOKE R/AU
L60
            923 S E3-E24, E43, E44
                 E MATUSKA M/AU
L61
             11 S E3-E5
                E NABER N/AU
L62
             45 S E3-E9
           1399 S L58-L62
L63
L64
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L67
          20077 S FLUORESCEIN/BI
           5771 S L1
L68
L69
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L70
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AN 2001-071424 [08] WPIX

DNN N2001-054040 DNC C2001-020049

Assaying mitochondrial membrane potential with energy transfer donor and acceptor molecules exogenous to the mitochondria, useful for identifying membrane potential modulating agents which are useful for treating diabetes and stroke.

DC B04 D16 S03

IN DYKENS, J A; GHOSH, S S; VELICELEBI, G

PA (MITO-N) MITOKOR

CYC 94

PI WO 2000079274 A2 20001228 (200108)* EN 189p G01N033-50

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TZ UG ZW

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AU 2000057636 A 20010109 (200122)

G01N033-50 C12P013-14

US 6280981 B1 20010828 (200151) C12

ADT WO 2000079274 A2 WO 2000-US17380 20000622; AU 2000057636 A AU 2000-57636 20000622; US 6280981 B1 Div ex US 1999-338122 19990622, US 2000-514569 20000223

FDT AU 2000057636 A Based on WO 200079274

PRAI US 2000-176383 20000114; US 1999-140433 19990622; US 1999-338122 19990622; US 2000-514569 20000223

IC ICM C12P013-14; G01N033-50

AB WO 200079274 A UPAB: 20010207

NOVELTY - A new method (M1) for assaying mitochondrial membrane potential comprises contacting a mitochondrial sample with energy transfer donor and energy transfer acceptor molecules exogenous to the mitochondria, exciting the donor molecule and detecting a signal generated by energy transfer between the donor and acceptor molecules.

DETAILED DESCRIPTION - A new method (M1) for assaying mitochondrial membrane potential comprises contacting a mitochondrial sample with energy transfer donor and energy transfer acceptor molecules exogenous to the mitochondria, exciting the donor molecule and detecting a signal generated by energy transfer between the donor and acceptor molecules.

In detail, M1 comprises:

- (a) contacting a sample comprising one or more mitochondria, simultaneously or sequentially and in either order, with each of a first and a second energy transfer molecule that is not endogenous to the mitochondria, where:
- (i) the first and second energy transfer molecules each localize independently of one another to the same submitochondrial site or to acceptably adjacent submitochondrial sites, the sites being selected from the mitochondrial outer membrane, mitochondrial inner membrane,



mitochondrial intermembrane space or mitochondrial matrix; and

- (ii) the first energy transfer molecule is an energy donor molecule and the second energy transfer molecule is an energy acceptor molecule;
- (b) exciting the energy donor molecule to produce an excited energy donor molecule; and
- (c) detecting a signal generated by energy transfer from the first energy transfer molecule to the second energy transfer molecule, where the concentration of at least one of the energy transfer molecules in the mitochondria changes as a function of membrane potential.

INDEPENDENT CLAIMS are also included for the following:

- (1) a method (M2) for identifying an agent that alters mitochondrial membrane potential, comprising:
- (a) steps (a) to (c) of M1, where step (a) is carried out in the presence or absence of the test compound; and
- (b) comparing the signal generated in the absence of the candidate agent to the signal generated in the presence of the candidate agent, and therefore identifying an agent that alters mitochondrial membrane potential;
- (2) a method (M3) for identifying a regulator of an agent that alters mitochondrial membrane potential, comprising:
- (a) steps (a) to (c) of M1, where step (a) is carried out in the presence or absence of the candidate regulator, and an agent that alters mitochondrial membrane potential or an agent identified by M2; and
- (b) comparing the signal generated in the absence of the candidate regulator to the signal generated in the presence of the candidate regulator, and therefore identifying a regulator that alters mitochondrial membrane potential;
- (3) a method (M4) for identifying an agent that preferentially alters mitochondrial membrane potential in mitochondria from a first biological source without substantially altering mitochondrial membrane potential in mitochondria from a second biological source;
- (4) a method (M5) of detecting the fusion of a first mitochondrion and a second mitochondrion;
- (5) a method (M6) of identifying an agent that alters the fusion of mitochondria;
- (6) a reagent for measuring mitochondrial Delta psi, comprising a FRET (fluorescence resonance energy transfer) donor molecule and a FRET acceptor molecule, where the accumulation of at least one of the molecules in mitochondria is dependent on Delta psi and the accumulation of the other molecules in mitochondria is independent of Delta psi;
- (7) a kit comprising the reagent of (6) and ancillary reagents for measuring mitochondrial Delta psi;
- (8) a method (M7) for assaying cellular membrane potential, comprising:
- (a) steps (a) and (b) of M1, where the sample comprises at least one cellular membrane instead of the mitochondria and the first and second energy transfer molecules each localize independently of one another to the same membrane site or to acceptably adjacent membrane sites such that at least one of the energy transfer molecules localizes to a cellular membrane that forms a subcellular compartment; and
- (b) detecting a signal generated by energy transfer from the first energy transfer molecule to the second energy transfer molecule, where the concentration of at least one of the energy transfer molecules in the membrane site changes as a function of membrane potential;
- (9) a method (M8) for identifying an agent that alters a cellular membrane potential;
- (10) a method (M9) for identifying a regulator of an agent that alters cellular membrane potential;
- (11) a method (M10) for identifying an agent that preferentially alters a cellular membrane potential in a membrane from a first biological source (or sample) without substantially altering cellular membrane potential in a membrane from a second biological source (or sample);
- (12) a method (M11) for detecting a specific type of cell in a sample, comprising:
 - (a) steps (a) and (b) of M1; and
 - (b) detecting a signal generated by energy transfer from the first



energy transfer molecule to the second energy transfer molecule, where at least one of the energy transfer molecules preferentially accumulates in the specific type of cell and the signal correlates with the presence of the specific type of cell in the sample;

- (13) a method (M12) for identifying a Delta psi m stabilizing agent;
- (14) a Delta psi m stabilizing agent identified by M12; and
- (15) a method (M13) of treating stroke comprising administering the Delta psi m stabilizing agent of (14) to a patient.

ACTIVITY - Nootropic; neuroprotective; antiparkinsonian; cytostatic; antipsoriatic; neuroleptic; cerebroprotective.

No biological data given.

MECHANISM OF ACTION - Mitochondrial membrane potential agonists and antagonists.

No biological data given.

USE - The method is used to develop assays of subcellular conditions or intracellular processes that are associated with diseases or disorders, e.g. Alzheimer's disease, Parkinson's disease or type II diabetes. The Delta psi m stabilizing agent is useful for treating stroke (all claimed).

Agents that alters a mitochondrial or cellular membrane potential are useful for treating diabetes, Alzheimer's disease, Parkinson's disease, schizophrenia, stroke, hyperproliferative diseases such as cancer and psoriasis.

The methods are also useful for to identify and characterize such agents.

Dwg.0/24

CPI EPI FS

FAAB; DCN

CPI: B04-B04H; B04-C01; B04-E01; B04-N0400E; B11-C07B3; B11-C08E; MC

B12-K04A; B12-K04E; B14-F01; B14-H01; B14-J01A3; B14-J01A4;

B14-J01B3; B14-L01; B14-L06; B14-N16; B14-N17C; B14-S04; D05-H09

EPI: S03-E14H

UPTX: 20010207 TECH

> TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Method: The excited energy donor molecule transfers energy to the energy acceptor molecule to produce an excited energy acceptor molecule, and the signal detected in step (c) results from energy released by the excited energy acceptor molecule. The energy transfer from the first energy transfer molecule to the second energy transfer molecule results in a decrease in the detectable signal. M1 further comprises contacting the mitochondria with an agent (i.e. an ionophore) that induces dissipation of mitochondrial membrane potential or an agent (e.g. CCCP (carbonyl cynaide m-chlorophenyl-hydrazone) and FCCP (carbonyl cyanide p-(trifluoromethoxy)phenyl-hydrazone)) that induces collapse of mitochondrial membrane potential. The sample is washed prior to the step of detecting a signal. The signal detected in step (c) is compared with a reference signal. The reference signal is generated by an indicator selected from an indicator of cell number, an indicator of mitochondrial mass, an indicator of cellular protein, an indicator of cellular DNA, an indicator of mitochondrial DNA, an indicator of mitochondrial protein or an indicator of fluid volume. The sample comprises one or more mitochondria that are present within at least one cell, and where the signal detected in step (c) is compared with a reference signal. The reference signal is generated from a subcellular site selected from a mitochondrial outer membrane, a mitochondrial inner membrane, a mitochondrial intermembrane space, a mitochondrial matrix, cytoplasm, nucleus, nuclear membrane or plasma membrane. Alternatively, the reference signal is generated from extracellular medium. The mitochondria are present within at least one cell during at least one step. The cell is an organism, a cultured cell, a cybrid cell, a plant cell or an animal cell.

The cell is present in a biological sample derived from a multicellular organism such as a plant or an animal such as a human. The human has, is suspected of having or is at risk of having a disease or disorder associated with organellar dysfunction, e.g. organellar dysfunction is mitochondrial dysfunction such as lysosomal dysfunction.

The first and second energy transfer molecules localize to a

submitochondrial site selected from the mitochondrial matrix or mitochondrial inner membrane. The concentration of the first energy transfer molecule in the submitochondrial site does not change as a function of membrane potential, and the concentration of the second energy transfer molecule in the mitochondrial matrix decreases as a function of membrane potential.

The first energy transfer molecule (F1) has an excitation maximum at a wavelength of 373 nm to 390 nm, and an emission maximum at a wavelength of 400 nm to 500 nm and the second energy transfer molecule (S1) has an excitation maximum at a wavelength of 400 nm to 500 nm. F1 is a fusion protein comprising:

- (a) a blue-shifted green fluorescent protein having a mutation in at least one of Phe-64, Ser-65, Tyr-66, Val-68 or Tyr-145; and
- (b) a polypeptide sequence that localizes the fusion protein to a submitochondrial site.
- S1 is selected from DASPEI (2-(4-(dimethylamino)styryl)-N-ethylpyridinium iodide), DASPMI (dimethylaminostyrylmethylpyridinium iodide), 4-Di-1-ASP (4-(4-(dimethylamino)styryl)-N-methylpyridinium iodide), 2-Di-1-ASP ASP (2-(4-(dimethylamino)styryl)-N-methylpyridinium iodide), DiOC7(3)
- (3,3'-diheptyloxadicarbocyanine iodide), DiOC6(3) (3,3'-
- dihexyloxadicarbocyanine iodide), JC-1 (5,5',6,6'-tetrachloro-1,1',3,3'-tetraethylbenzimidazoylcarbocyanine iodide) and SYTO (RTM) 18 yeast mitochondrial stain.

The first energy transfer molecule (F2) has an excitation maximum at a wavelength of 425 nm to 440 nm, and an emission maximum at a wavelength of 450 nm to 535 nm and the second energy transfer molecule (S2) has an excitation maximum at a wavelength of 450 nm to 530 nm. F2 is a fusion protein comprising:

- (a) a cyan-shifted green fluorescent protein polypeptide having a mutation in at least one of Phe-64, Ser-65, Tyr-66, Asn-146, Met-153, Val-163 or Asn-212; and
- (b) a polypeptide sequence that localizes the fusion protein to a submitochondrial site.
- S2 is selected from DASPEI, 2-Di-1-ASP, DiOC6(3), SYTO (RTM) 18 yeast mitochondrial stain, rhodamine 6G, JC-1, NBD C6-ceramide or NBD C6-sphingomyelin

The first energy transfer molecule (F3) has an excitation maximum at a wavelength of 470 mn to 500 nm, and an emission maximum at a wavelength of 505 nm to 565 nm and the second energy transfer molecule (S3) has an excitation maximum at a wavelength of 505 nm to 565 nm. F3 is selected from nonylacridine orange (NAO), MitoTracker (RTM) Green FM, MitoFluor (RTM) Green, or a fusion protein, where the fusion protein comprises:

- (a) a green fluorescent protein selected from a wildtype green fluorescent protein, a red-shifted green fluorescent protein having a mutation in one or more of Phe-64, Ser-65, Tyr-66, Gln-69, Ser-72 and Thr-203, or a yellow-shifted green fluorescent protein having a mutation in one or more of Phe-64, Ser-65, Tyr-66, Gln-69, Ser-72 or Thr-203; and
- (b) polypeptide sequence that localizes the fusion protein to a submitochondrial site.

S3 is selected from rhodamine 123, JC-1, tetrabromorhodamine 123, rhodamine 6G, TMRM (tetramethylrhodamine, methyl ester), TMRE (tetramethylrhodamine, ethyl ester), tetramethylrosamine or rhodamine B. The first energy transfer molecule (F4) has an excitation maximum at a wavelength of 545 mn to 560 nm, and an emission maximum at a wavelength of 565 nm to 625 nm and the second energy transfer molecule (S4) has an excitation maximum at a wavelength of 565 nm to 625 nm. F4 is selected from MitoTracker (RTM) Orange CMTMRos and S4 is DiOC2(5) (3,3'-diethyloxadicarbocyanine iodide).

The first energy transfer molecule (F5) has an excitation maximum at a wavelength of 495 mn to 510 nm, and an emission maximum at a wavelength of 510 nm to 570 nm and the second energy transfer molecule (S5) has an excitation maximum at a wavelength of 510 nm to 560 nm. F5 is selected from a fusion protein comprising:

(a) a polypeptide sequence selected from 'FLASH' (fluorescein arsenical helix binder) protein or a yellow-shifted green fluorescent protein sequence having a mutation in one or more of

Ser-65, Tyr-66, Ser-72 or Thr-203; and

- (b) a polypeptide sequence that localizes the fusion protein to a submitochondrial site.
- S5 is selected from JC-1, tetrabromorhodamine 123, rhodamine 6G, TMRM, TMRE, tetramethylrosamine, rhodamine B or 4-dimethylamino-tetramethylrosamine.
- A relative amount of the signal generated by energy transfer is detected. The signal is detected over a period of time and integrated, and a rate of change in the signal level is determined. The membrane potential comprises an electric potential, a pH potential, or both. The first and second energy transfer molecules localize to within 10 to 100, preferably 20 to 50, angstroms of each other. The signal is generated by fluorescence resonance energy transfer.
- In M3, the regulator is an agonist or antagonist of the agent that alters mitochondrial potential. The agent is an apoptogen, a thapsigargin, an ionophore (e.g. ionomycin or A23187), or an excitatory amino acid (e.g. glutamate, NAAG (undefined), NMDA (N-methyl-D-aspartate), AMPA (undefined), APPA (undefined) or kainate) or its derivatives.

 M4 comprises:
- (a) contacting, in the absence and presence of a candidate agent, a biological sample (from each biological source) comprising one or more mitochondria simultaneously or sequentially and in either order with a first and a second energy transfer molecule that is not endogenous to the mitochondria;
- (b) steps (a)(i), (a)(ii), (b) and (c) of M1; and
- (c) comparing the signal generated in each sample in the absence of the candidate agent to the signal generated in each sample in the presence of the candidate agent, and therefore identifying an agent that preferentially alters mitochondrial membrane potential.
- The first and second biological samples are from distinct biological sources, preferably tissues. The first biological source is a mammal suspected of having, diagnosed as having or predisposed to having a disease, and the second biological source is a mammal that is not suspected of having and has not been diagnosed as having or predisposed to having the disease. The first and second biological sources are both human. The disease is Alzheimer's disease, Parkinson's disease or type II diabetes. When the biological source is a tissue, the first and second tissues are derived from the same subject, a subject of the same species or subjects of distinct species.

M5 comprises:

- (a) contacting a first sample comprising one or more mitochondria with a first energy transfer molecule that is not endogenous to the mitochondria; (b) contacting a second sample comprising one or more mitochondria with a second energy transfer molecule that is not endogenous to the mitochondria, where the first energy transfer molecule is an energy donor molecule and the second energy transfer molecule is an energy acceptor molecule;
- (c) contacting the first sample with the second sample under conditions and for a time sufficient to permit mitochondrial fusion;
- (d) exciting the energy donor molecule to produce an excited energy donor molecule; and
- (e) detecting a signal generated by energy transfer from the first energy transfer molecule to the second energy transfer molecule, and therefore determining fusion of the first mitochondrion and the second mitochondrion;

M6 comprises:

- (a) steps (a) and (b) of M5;
- (b) carrying out step (c) of M5 in the presence and absence of the candidate agent;
- (c) steps (d) of M5;
- (d) detecting a signal generated by energy transfer from the first energy transfer molecule to the second energy transfer molecule; and
- (e) comparing the signal detected in the absence of the candidate agent to the signal detected in the presence of the candidate agent, and therefore identifying an agent that alters the fusion of the mitochondria.
- In M5 and M6, the first and second energy transfer molecules have the

properties described in (a)(i) and (a)(ii) or M1.

In M2 to M5, the agent increases, dissipates or collapses mitochondrial membrane potential, or alters an equilibrium distribution of at least one ionic species (e.g. calcium) on either side of a cellular membrane (e.g. mitochondrial membrane). The agent (A1) that collapses mitochondrial membrane potential is an apoptogen and it interacts with an adenine nucleotide translocator. A1 is an atractyloside, carboxyatractyloside, bongkrekic acid or isobongkrekic acid.

In M7, the first energy transfer molecule localizes to a first membrane site selected from mitochondria, endoplasmic reticulum, golgi, lysosome or plasma membrane and the second energy transfer molecule localizes to the same membrane site or to an acceptably adjacent membrane site selected from mitochondria, endoplasmic reticulum, golgi, lysosome or plasma membrane. The concentration of the first energy transfer molecule in the first membrane site does not change as a function of membrane potential, and the concentration of the second energy transfer molecule in the membrane site decreases as a function of membrane potential. The first energy transfer molecule is F1, F2, F3 or F4 and the second energy transfer molecule is S1, S2, S3 or S4, respectively.

M8 comprises:

- (a) contacting, in the absence and presence of a candidate agent, a sample comprising one or more cellular membranes simultaneously or sequentially and in either order with each of a first and a second energy transfer molecule that is not endogenous to the sample, where:
- (i) the first and second energy transfer molecules each localize independently of one another to the same membrane site or to acceptably adjacent membrane sites such that at least one of the energy transfer molecules localizes to a cellular membrane that forms a subcellular compartment, and
- (ii) the first energy transfer molecule is an energy donor molecule and the second energy transfer molecule is an energy acceptor molecule;
- (b) exciting the energy donor molecule to produce an excited energy donor molecule;
- (c) detecting a signal generated by energy transfer from the first energy transfer molecule to the second energy transfer molecule, where the concentration of at least one of the energy transfer molecules in the subcellular compartment changes as a function of membrane potential; and (d) comparing the signal generated in the absence of the candidate agent
- (d) comparing the signal generated in the absence of the candidate agent to the signal generated in the presence of the candidate agent, and therefore identifying an agent that alters cellular membrane potential. M9 comprises:
- (a) steps (a) to (c) of M8, where step (a) is carried out in the presence or absence of the candidate regulator, and an agent that alters cellular membrane potential or an agent identified by M8; and
- (b) comparing the signal generated in the absence of the candidate regulator to the signal generated in the presence of the candidate regulator, and therefore identifying a regulator that alters mitochondrial membrane potential.
- M10 comprises steps (a) and (b) of M4, where the biological sample (from each biological source) comprising one or more cellular membranes instead of one or more mitochondria. Step (c) comparing the signal generated in each sample in the absence of the candidate agent to the signal generated in each sample in the presence of the candidate agent, and therefore identifying an agent that preferentially alters cellular membrane potential. The first and second biological samples are from distinct biological sources, preferably tissues.
- Mll further comprises comparing the signal generated in the sample with the signal generated from a control sample lacking the specific type of cell. The specific type of cell is a cancer cell Ml2 comprises:
- (a) contacting, in the absence and presence of a candidate DELTA(psi)m stabilizing agent, an agent that alters DELTA(psi)m and a sample comprising one or more mitochondria simultaneously or sequentially and in either order with each of a first and a second energy transfer molecule that is not endogenous to the mitochondria, where the energy transfer molecules have the properties as described in (a)(i) and (a)(ii) of M1;

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(b) steps (b) and (c) of M1;
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(c) comparing the signal generated in the absence of the candidate DELTA(psi)m stabilizing agent, to the signal generated in the presence of the candidate DELTA(psi)m stabilizing agent, and therefore identifying DELTA(psi)m stabilizing agent.

The mitochondria are contained within cells. The agent that alters DELTA(psi)m is an agent that increases the level of cytosolic Ca2+. The agent that increases the level of cytosolic Ca2+ is selected from calcium ionophore or thapsigargin. The cells comprise one or more types of glutamate receptors. Alternatively, the agent that increases the level of cytosolic Ca2+ is an excitatory amino acid or its derivative, e.g. glutamate, NAAG, NMDA, AMPA, APPA and kainate.

In M1, M12 and M13, the cell is a permeabilized cell.

Preferred Reagent: The molecule that accumulates in mitochondria independent of DELTA(psi) is selected from NAO, MitoTracker (RTM) Green FM, MitoFluor (RTM), DAPI (4', 6-diamino-2-phenylindole), and a fusion protein comprising:

(a) a polypeptide selected from a red-shifted green fluorescent protein, a yellow-shifted green fluorescent protein and a 'FLASH' polypeptide, and (b) a polypeptide sequence that localizes the fusion protein to the mitochondrial matrix or inner membrane.

The molecule that accumulates in mitochondria in a manner dependent on DELTA(psi) is selected of TMRM, TMRE, rhodamine 123, ethidum bromide, 4-Di-1-ASP, 2-Di-1-ASP or DASPEI.

The first FRET molecule that accumulates in mitochondria is formulated to dissolve to an extent necessary to saturate a population of cells in an aqueous solution with the first molecule within 0.01 to 2 minutes after being contacted with it, and the second molecule that accumulates in mitochondria is formulated to dissolve to an extent necessary to saturate a population of cells in an aqueous solution with the second molecule within 2.5 to 5 minutes after being contacted with it. One of the molecules that accumulates in mitochondria is dissolved in an aqueous solution, and the other of the molecules that accumulates in mitochondria is present in solid form in the reagent. The molecule that accumulates in mitochondria and that is present in solid form in the reagent is formulated to dissolve to an extent necessary to saturate a population of cells in an aqueous solution with the second molecule within 0.01 to 5 minutes after being contacted with it.

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L94 ANSWER 2 OF 2 WPIX COPYRIGHT 2001 DERWENT INFORMATION LTD
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AN 1999-288410 [24] WPIX

DNN N1999-215341 DNC C1999-085363

TI Biarsenical compounds that react specifically with cysteine residues.

DC B04 B05 D16 S03

IN GRIFFIN, B A; TSIEN, R Y; GRIFFIN, A B

PA (REGC) UNIV CALIFORNIA

CYC 83

PI WO 9921013 A1 19990429 (199924)* EN 76p G01N033-566 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SZ UG ZW

W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE GH GM HR HU ID IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG US UZ VN YU ZW

US 5932474 A 19990803 (199937) C12N015-63 AU 9911139 A 19990510 (199938) G01N033-566 US 6008378 A 19991228 (200007) C07F009-80 US 6054271 A 20000425 (200027) G01N033-566 EP 1032837 A1 20000906 (200044) EN G01N033-566

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE ADT WO 9921013 A1 WO 1998-US22363 19981021; US 5932474 A US 1997-955206 19971021; AU 9911139 A AU 1999-11139 19981021; US 6008378 A US 1997-955859 19971021; US 6054271 A US 1997-955050 19971021; EP 1032837 A1 EP 1998-953881 19981021, WO 1998-US22363 19981021

FDT AU 9911139 A Based on WO 9921013; EP 1032837 Al Based on WO 9921013

```
PRAI US 1997-955859 19971021; US 1997-955050
                                                 19971021; US 1997-955206
     19971021
IC
     ICM C07F009-80; C12N015-63; G01N033-566
     ICS C07D493-10; C12N015-09; C12N015-64
AB
     WO 9921013 A UPAB: 19990624
     NOVELTY - Biarsenical compounds (BC) able to react specifically
     with cysteine residues in a target sequence to generate a detectable
     signal are new.
          DETAILED DESCRIPTION - (BC) have formula (I) including tautomers,
     anhydrides and salts:
          X1 and X2 = chloro, bromo, iodo, ORa or SRa., or both together form
     groups of formula (f1)-(f4);
          Ra = hydrogen, 1-4C alkyl, 2-hydroxyethyl, carboxymethyl or cyano;
          Z = 1,2-ethanediyl, 1,2- or 1,3-propanediyl, 2,3-butanediyl,
     1,2-phenylene (optionally 4-methyl substituted), 1,2-cycopentanediyl,
     1,2-cyclohexanediyl, 3-(hydroxy or sulfo)-1,2-propanediyl or
     1,2-bis(carboxy)-1,2-ethanediyl;
          Y1 and Y2 = hydrogen or methyl, or together form a ring such that
     (I) has the formula (I')
         M = oxygen, sulfur, methylene, dimethylmethylene or imino;
          R1 and R2 = ORa, acetyloxy, NRaRb or hydrogen;
          R3 and R4 = hydrogen, fluoro, chloro, bromo, iodo, ORa or Ra;
          R1 and R2, and/or R3 and R4 together form a ring in which:
          (i) one of R1 or R3 is 2-3C alkyl and the other NRa, and
          (ii) one of R2 and R4 is 2-3C alkyl and the other is NRa;
     Rb is as Ra;
          Q = CRaRb, CRaORb, CO or a spirolactone of formulae (Ia), (Ib) and
     (Ic) with the spiro linkage at C1:
          INDEPENDENT CLAIMS are also included for the following:
          (1) a binding partner (BP) comprising carrier polypeptide (CP) and
    heterologous target sequence (TS) which contains at least one Cys that can
    react specifically with the compound (II);
          (2) vector containing a nucleic acid sequence that encodes BP;
          (3) host cell containing an exogenous BP;
          (4) method for labeling a carrier by reacting BP with (I);
          (5) crosslinking two BP by reacton with a tetra-arsenical
    compound (tac);
          (6) kits containing (BC) plus BP or a vector encoding TS;
          (7) complex of (BC) and TS; and
          (8) (tac) consisting of two (BC) coupled through a linking group:
         ACTIVITY - None given.
         MECHANISM OF ACTION - None given.
         USE - (I) are used:
          (i) as labels that allow identification of carrier molecules, e.g. in
    polypeptide purification, immunoassays or other chemical or biological
    assays, including labeling in vivo, e.g. to identify, locate or quantify
    polypeptides or nucleic acids);
          (ii) for attaching a polypeptide to a solid substrate; or
          (iii) to induce a polypeptide domain to adopt a more nearly alpha -
    helical form, e.g. a conformation that can bind a drug.
          Tetra-arsenical compounds derived from (I) are used to crosslink two
    binding partners, e.g. to study the effect of dimerization on signal
     transduction.
         ADVANTAGE - (I) react specifically with Cys-containing targets, and
    can be engineered to have particular properties, especailly ability to
    cross a biological membrane and absence of any self-fluorescence. Both (I)
    and its target sequence are small, and (I) binding between them is
    reversible, e.g. by treatment with a dithiol. Particularly (I) becomes
    fluorescent when bound to its target, but with a significant red-shift
    from the fluorescence of fluorescein, allowing detection with very low
    background.
    CPI EPI
FS
    AB; GI; DCN
FA
    CPI: B12-K04A; D05-H09; D05-H10; D05-H12E; D05-H14
MC
     EPI: S03-E14H4
```

TECH

UPTX: 19990624

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preferred binding partners: BP has a TS attached to either end of CP, or internally. TS preferably has four Cys, especially the alpha-helical domain Cys-Cys-X-Y-Cys-Cys X and Y = amino acids, preferably those with high alpha-helical propensity

. CP is particularly an antibody or enzyme.

Preferred compounds: In (I), Xl and X2 particularly form SCH2CH2S and Q is a spiro-lactone. (I) are particularly able to cross a biological membrane and may be substituted by one or more detectable groups. Optionally it is immobilized on a solid phase.

Preparation: Typically, fluorescein mercuric acetate (commercially available) was reacted with arsenic trichloride in presence of palladium diacetate, and the resulting bis-dicholorarsino derivative reacted with a dithiol. (tac) are produced similarly from a tetrakis (acetomercuri) bifluorescein.

Process: To label a carrier molecule, BP is reacted with (I). Optionally (I) or TS is immobilized and (I) may subsequently released from TS. The signal, particularly fluorescent, generated by (I) may be monitored. The crosslinking reaction of (e) may involve same or different BP.

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred vectors: The vectors include a gene containing coding sequences for both TS and CP.

Preferred host cells: The cells are bacteria, yeast, insect, mammalian or plant cells.

=> d his 182-

(FILE 'BIOSIS' ENTERED AT 16:04:07 ON 13 NOV 2001)

```
FILE 'WPIX' ENTERED AT 16:05:09 ON 13 NOV 2001
L82
           1394 S L67
                E FLUORESCEIN/DCN
                E E3+ALL
            643 S E2 OR 1594/DRN
L83
L84
             51 S E4
             11 S E6
L85
L86
           1792 S L82~L85
L87
             14 S L86 AND ARSEN?
             10 S L86 AND (B133 OR B233 OR B333 OR B433 OR B533 OR B633)/M0,M1,
F88
L89
             16 S L87, L88
             14 S L86 AND ?ARSEN?
L90
L91
             16 S L89, L90
L92
              2 S L91 AND HELI?
              1 S L91 AND (BISARSEN? OR BIARSEN? OR BI ARSEN?)
L93
L94
              2 S L92, L93
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FILE 'WPIX' ENTERED AT 16:09:35 ON 13 NOV 2001 SET COST ON

=> file medline, biosis, toxlit, embase, dgene, uspat, wpids, japio, jicst, frosti, fsta, cen, ceaba, biotechds, scisearch, agricola

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.30 0.30

FULL ESTIMATED COST

FILE 'MEDLINE' ENTERED AT 15:07:01 ON 02 NOV 2001

FILE 'BIOSIS' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 BIOSIS(R)

FILE 'TOXLIT' ENTERED AT 15:07:01 ON 02 NOV 2001

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FILE 'DGENE' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 DERWENT INFORMATION LTD

FILE 'USPATFULL' ENTERED AT 15:07:01 ON 02 NOV 2001
CA INDEXING COPYRIGHT (C) 2001 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'WPIDS' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 DERWENT INFORMATION LTD

FILE 'JAPIO' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 Japanese Patent Office (JPO)

FILE 'JICST-EPLUS' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 Japan Science and Technology Corporation (JST)

FILE 'FROSTI' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 Leatherhead Food Research Association

FILE 'FSTA' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 International Food Information Service

FILE 'CEN' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 American Chemical Society (ACS)

FILE 'CEABA-VTB' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (c) 2001 DECHEMA eV

FILE 'BIOTECHDS' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 DERWENT INFORMATION LTD

FILE 'SCISEARCH' ENTERED AT 15:07:01 ON 02 NOV 2001 COPYRIGHT (C) 2001 Institute for Scientific Information (ISI) (R)

FILE 'AGRICOLA' ENTERED AT 15:07:01 ON 02 NOV 2001

=> s fluorescein arsenical helix binder or FLAsh

L1 188451 FLUORESCEIN ARSENICAL HELIX BINDER OR FLASH

=> s 11 and modified

L2 29523 L1 AND MODIFIED

=> s 12 and target sequence motif

```
11 FILES SEARCHED...
             0 L2 AND GET SEQUENCE MOTIF
L3
=> s 12 and target sequence
           467 L2 AND TARGET SEQUENCE
L4
=> s polypeptide () method () isolation
  11 FILES SEARCHED...
             O POLYPEPTIDE (W) METHOD (W) ISOLATION
L5
=> s polypeptide and method of isolated
  10 FILES SEARCHED...
L6
           209 POLYPEPTIDE AND METHOD OF ISOLATED
=> s 16 and 14
             1 L6 AND L4
L7
=> d 17 ti abs ibib tot
L7
     ANSWER 1 OF 1 USPATFULL
       Thermophilic polymerase III holoenzyme
{f TI}
       The present invention relates to gene and amino acid sequences encoding
AB
       DNA polymerase III holoenzyme subunits and structural genes from
       thermophilic organisms. In particular, the present invention provides
       DNA polymerase III holoenzyme subunits of T. thermophilus. The present
       invention also provides antibodies and other reagents useful to
identify
       DNA polymerase III molecules.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
                        2001:78932 USPATFULL
ACCESSION NUMBER:
                        Thermophilic polymerase III holoenzyme
TITLE:
                        McHenry, Charles S., Denver, CO, United States
INVENTOR(S):
                        Seville, Mark, Denver, CO, United States
                        Cull, Millard G., Denver, CO, United States
                        University Technology Corporation, CO, United States
PATENT ASSIGNEE(S):
                        (U.S. corporation)
                             NUMBER
                        US 6238905 B1 20010529
US 1997-928213 19970912
PATENT INFORMATION:
                                                19970912 (8)
APPLICATION INFO.:
DOCUMENT TYPE:
                        Utility
FILE SEGMENT:
                        Granted
PRIMARY EXAMINER:
                        Stole, Einar
                        Medlen & Carroll, LLP
LEGAL REPRESENTATIVE:
NUMBER OF CLAIMS:
                        39
EXEMPLARY CLAIM:
                        1
NUMBER OF DRAWINGS:
                        29 Drawing Figure(s); 18 Drawing Page(s)
LINE COUNT:
                        4725
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
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=> d his

(FILE 'HOME' ENTERED AT 15:06:11 ON 02 NOV 2001)

FILE 'MEDLINE, BIOSIS, TOXLIT, EMBASE, DGENE, USPATFULL, WPIDS, JAPIO, JICST-EPLUS, FROSTE FSTA, CEN, CEABA-VTB, BIOTECHT SCISEARCH, AGRICOLA' ENTERED 15:07:01 ON 02 NOV 2001

L1 188451 S FLUORESCEIN ARSENICAL HELIX BINDER OR FLASH

L2 29523 S L1 AND MODIFIED

L3 0 S L2 AND TARGET SEQUENCE MOTIF

L4 467 S L2 AND TARGET SEQUENCE

L5 0 S POLYPEPTIDE () METHOD () ISOLATION
L6 209 S POLYPEPTIDE AND METHOD OF ISOLATED

L7 1 S L6 AND L4

=> s 14 and (polypeptide)

L8 139 L4 AND (POLYPEPTIDE)

=> s 18 and isolation

L9 101 L8 AND ISOLATION

=> s 19 and acylation

L10 46 L9 AND ACYLATION

=> s 110 and alanine

L11 38 L10 AND ALANINE

=> s 111 and agarose

L12 36 L11 AND AGAROSE

=> d 112 ti abs ibib tot

L12 ANSWER 1 OF 36 USPATFULL

TI Human galectin homolog

AB The present invention provides a polynucleotide which identifies and encodes a novel human galectin-8. The invention provides for genetically

engineered expression vectors and host cells comprising the nucleic acid

sequence encoding human galectin-8. The invention also provides for the production and use of substantially purified human galectin-8 in pharmaceutical compositions to increase immune responses. The invention also provides for the use of antisense molecules and antibodies in pharmaceutical compositions to decrease immune response. The invention also describes diagnostic assays which utilize the polynucleotide to hybridize with the transcripts and/or genomic DNA encoding human galectin-8 and anti-human galectin-8 antibodies which specifically bind to human galectin-8.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:142466 USPATFULL TITLE: Human galectin homolog

INVENTOR(S): Hawkins, Phillip R., Mountain View, CA, United States

Bandman, Olga, Mountain View, CA, United States

PATENT ASSIGNEE(S): Incyte Genomics, Inc., Palo Alto, CA, United States

(U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6281333 B1 20010828

APPLICATION INFO.: US 1998-212146 19981215 (9)

RELATED APPLN. INFO.: Division of Ser. No. US 1996-728521, filed on 9 Oct

1996, now patented, Pat. No. US 5869289

DOCUMENT TYPE: Itility FILE SEGMENT: RANTED

PRIMARY EXAMINER: Achutamurthy, Ponnathapu

ASSISTANT EXAMINER: Tung, Peter P.

LEGAL REPRESENTATIVE: Incyte Genomics, Inc.

NUMBER OF CLAIMS: 4
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 4 Drawing Figure(s); 4 Drawing Page(s)

LINE COUNT: 2410

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 2 OF 36 USPATFULL

TI Human KDEL receptor

The present invention provides a novel human KDEL receptor (NHKR) and polynucleotides which identify and encode NHKR. The invention also provides genetically engineered expression vectors and host cells comprising the nucleic acid sequences encoding NHKR and a method for producing NHKR. The invention also provides for agonists, antibodies,

or

antagonists specifically binding NHKR, and their use, in the prevention and treatment of diseases associated with expression of NHKR. Additionally, the invention provides for the use of antisense molecules to polynucleotides encoding NHKR for the treatment of diseases associated with the expression of NHKR. The invention also provides diagnostic assays which utilize the polynucleotide, or fragments or the complement thereof, and antibodies specifically binding NHKR.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:106057 USPATFULL TITLE: Human KDEL receptor

INVENTOR(S): Bandman, Olga, Mountain View, CA, United States
Hillman, Jennifer L., San Jose, CA, United States

Goli, Surya K., Sunnyvale, CA, United States

19980813 (9)

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

PATENT INFORMATION: US 6103874 KIND DATE 20000815

US 1998-133735

RELATED APPLN. INFO.: Division of Ser. No. US 1996-753159, filed on 21 Nov

1996, now patented, Pat. No. US 5824500

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Kemmerer, Elizabeth ASSISTANT EXAMINER: Basi, Nirmal S.

LEGAL REPRESENTATIVE: Incyte Pharmaceuticals, Inc.

NUMBER OF CLAIMS: 5
EXEMPLARY CLAIM: 1

APPLICATION INFO.:

NUMBER OF DRAWINGS: 5 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 2107

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 3 OF 36 USPATFULL

TI Human ATP binding-cassette transport protein

The invention provides a human ATP-binding cassette transport protein (ABCtxH) and polynucleotides which identify and encode ABCtxH. The invention also provides expression vectors, host cells, agonists, antibodies and antagonists. The invention also provides methods for treating disorders associated with expression of ABCtxH.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:80850 USPATFULL

TITLE: Human ATP binding-cassette transport protein

INVENTOR(S): willman, Jennifer L., Mountain w, CA, United States

hah, Purvi, Sunnyvale, CA, United States

Corley, Neil C., Mountain View, CA, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

RELATED APPLN. INFO.: Division of Ser. No. US 1997-895522, filed on 17 Jul

1997, now patented, Pat. No. US 5858719

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Carlson, Karen Cochrane
LEGAL REPRESENTATIVE: Incyte Pharmaceuticals, Inc.

NUMBER OF CLAIMS: 6
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 13 Drawing Figure(s); 13 Drawing Page(s)

LINE COUNT: 2068

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 4 OF 36 USPATFULL

TI Human actVA-ORF4-like protein

The invention provides a human actVA-ORF4-like protein (A ORFP) and polynucleotides which identify and encode A-ORFP. The invention also provides expression vectors, host cells, agonists, antibodies and antagonists. The invention also provides methods for treating disorders associated with expression of A-ORFP.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:80731 USPATFULL

TITLE: Human actVA-ORF4-like protein

INVENTOR(S): Lal, Preeti, Santa Clara, CA, United States

Tang, Tom, San Jose, CA, United States

Corley, Neil C., Mountain View, CA, United States

Shah, Purvi, Sunnyvale, CA, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6080723 20000627 APPLICATION INFO.: US 1998-216294 19981218 (9

RELATED APPLN. INFO.: Division of Ser. No. US 1997-923856, filed on 3 Sep

1997, now patented, Pat. No. US 5928894

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Prouty, Rebecca E. ASSISTANT EXAMINER: Longton, Enrique D.

LEGAL REPRESENTATIVE: Hamlet-Cox, DianaIncyte Pharmaceuticals, Inc.

NUMBER OF CLAIMS: 5
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 7 Drawing Figure(s); 7 Drawing Page(s)

LINE COUNT: 2284

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 5 OF 36 USPATFULL

TI Cyclic nucleotide phosphodiesterases

AB The invention provides human cyclic nucleotide phosphodiesterases (PDE8)

and polynucleotides which identify and encode PDE8. The invention also provides expression vectors, host cells, antibodies, agonists, and

antagonists. The invention also provides methods for treating or preventing diso rs associated with expression PDE8.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:80556 USPATFULL

TITLE: Cyclic nucleotide phosphodiesterases

INVENTOR(S):

Au-Young, Janice, Berkeley, CA, United States

Cocks, Benjamin G, Palo Alto, CA, United States

Cocks, Benjamin G., Palo Alto, CA, United States Coleman, Roger, Mountain View, CA, United States Seilhamer, Jeffrey J., Los Altos, CA, United States

Fisher, Douglas A., Groton, CA, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

PATENT INFORMATION: US 6080548 KIND DATE 20000627

APPLICATION INFO.: US 1999-255748 19990223 (9)

RELATED APPLN. INFO.: Division of Ser. No. US 1997-974565, filed on 19 Nov

1997, now patented, Pat. No. US 5932423

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schwartzman, Robert A.

LEGAL REPRESENTATIVE: Incyte Pharmaceuticals, Inc, Murry, Lynn E.

NUMBER OF CLAIMS: 6
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 32 Drawing Figure(s); 32 Drawing Page(s)

LINE COUNT: 2831

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 6 OF 36 USPATFULL
TI Transducin beta-1 subunit

The invention provides a human transducin beta-1 subunit (TBS) and polynucleotides which identify and encode TBS. The invention also provides expression vectors, host cells, agonists, antibodies, and antagonists. The invention also provides methods for treating or preventing diseases associated with expression of TBS.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:77204 USPATFULL

TITLE: Transducin beta-1 subunit

INVENTOR(S): Bandman, Olga, Mountain View, CA, United States

Lal, Preeti, Santa Clara, CA, United States

Corley, Neil C., Mountain View, CA, United States

19971106 (8)

Shah, Purvi, Sunnyvale, CA, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

APPLICATION INFO.: US 1997-965600

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted

PRIMARY EXAMINER: Carlson, Karen Cochrane

ASSISTANT EXAMINER: Stole, Einar

LEGAL REPRESENTATIVE: Muenzen, Colette C. Incyte Pharmaceuticals, Inc.

NUMBER OF CLAIMS: 9
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 3 Drawing Figure(s); 7 Drawing Page(s)

LINE COUNT: 2122

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 7 OF 36 USPATFULL

TI G-protein coupled receptors associated with immuma response

The invention prides two human G-protein couple receptors associated with immune response (GRIR) and polynucleotides which identify and encode GRIR. The invention also provides expression vectors, host

cells,

antibodies, agonists, and antagonists. The invention also provides methods for treating or preventing disorders associated with expression of GRIR.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:61412 USPATFULL

TITLE: G-protein coupled receptors associated with immune

response

INVENTOR(S): Lal, Preeti, Santa Clara, CA, United States

Bandman, Olga, Mountain View, CA, United States

Hillman, Jennifer L., Mountain View, CA, United States

Yue, Henry, Sunnyvale, CA, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6063596 20000516

APPLICATION INFO.: US 1997-988876 19971211 (8)

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Ulm, John

LEGAL REPRESENTATIVE: Muenzen, Colette C.Incyte Pharmaceuticals, Inc.

NUMBER OF CLAIMS: 12
EXEMPLARY CLAIM: 1
NUMBER OF DRAWINGS: 15

NUMBER OF DRAWINGS: 15 Drawing Figure(s); 15 Drawing Page(s)

LINE COUNT: 2777

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 8 OF 36 USPATFULL

TI Human prostate-associated protease

The present invention provides a human prostate-associated protease (HUPAP) and polynucleotides which identify and encode HUPAP. The invention also provides expression vectors, host cells, antibodies, antagonists, and antisense molecules. The invention also provides methods for treating disorders associated with expression of HUPAP.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:37590 USPATFULL

TITLE: Human prostate-associated protease

INVENTOR(S): Bandman, Olga, Mountain View, CA, United States

Lal, Preeti, Sunnyvale, CA, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6043033 20000328
APPLICATION INFO.: US 1997-807151 19970227 (8)

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Wax, Robert A.

LEGAL REPRESENTATIVE: Murry, Lynn E. Incyte Pharmaceuticals, Inc.

NUMBER OF CLAIMS: 10
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 5 Drawing Figure(s); 5 Drawing Page(s)

LINE COUNT: 2114

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 9 OF 36 USPATFULL

TI Nucleic acid se nce of senescence associated e

Human gene GC6 — expressed more abundantly in senescent cells than young cells. Isolated, purified, and recombinant nucleic acids and proteins corresponding to the human GC6 gene and its mRNA and protein products, as well as peptides and antibodies corresponding to the GC6 protein can be used to identify senescent cells, distinguish between senescent and young cells, identify agents that alter senescent gene expression generally and GC6 expression specifically; such agents as well as GC6 gene and gene products and products corresponding thereto can be used to prevent and treat diseases and conditions relating to cell senescence.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:18280 USPATFULL

TITLE: Nucleic acid sequence of senescence asssociated gene

INVENTOR(S): Funk, Walter, Hayward, CA, United States

PATENT ASSIGNEE(S): Geron Corporation, Menlo Park, CA, United States (U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6025194 20000215

APPLICATION INFO.: US 1997-974180 19971119 (8)

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted

PRIMARY EXAMINER: Huff, Sheela

ASSISTANT EXAMINER: Bansal, Geetha P.

LEGAL REPRESENTATIVE: Earp, David J., Kaster, Kevin

NUMBER OF CLAIMS: 10
EXEMPLARY CLAIM: 1,6
LINE COUNT: 4667

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 10 OF 36 USPATFULL

TI Nucleic acids encoding human tyrosine phosphatases

AB The present invention provides novel human protein tyrosine phosphatases

(HPTP) and polynucleotides which identify and encode HPTP. The invention

provides for genetically engineered expression vectors and host cells comprising the nucleic acid sequences encoding HPTP and for a method of producing HPTP. The invention also provides for pharmaceutical compositions comprising HPTP or antagonists of HPTP, and antibodies which specifically bind HPTP. Additionally, the invention provides antisense molecules to HPTP for treatment or prevention of diseases associated with abnormal expression of HPTP.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:12642 USPATFULL

TITLE: Nucleic acids encoding human tyrosine phosphatases

INVENTOR(S): Goli, Surya K., Sunnyvale, CA, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Teng, Sally P.

LEGAL REPRESENTATIVE: Mohan-Peterson, Sheela, Billings, Lucy J.Incyte

Pharmaceuticals, Inc.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 11 Drawing Figure(s); 11 Drawing Page(s)

LINE COUNT: 1932

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 11 OF 36 USPATFULL

TI Human proteasome subunit proteins

AB The present invention provides polynucleotides which identify and encode

novel human proteasome subunit proteins. The invention provides for genetically engineered expression vectors and host cells comprising the nucleic acid sequences encoding PSUB. The invention also provides for the use of substantially purified PSUB, antagonists, and in pharmaceutical compositions for the treatment of diseases associated with the expression of PSUB. Additionally, the invention provides for the use of antisense molecules to PSUB in pharmaceutical compositions for treatment of diseases associated with the expression of PSUB. The invention also describes diagnostic assays which utilize diagnostic compositions comprising the polynucleotide, fragments or the complement thereof, which hybridize with the genomic sequence or the transcript of polynucleotides encoding PSUB or anti-PSUB antibodies which

specifically

bind to PSUB.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2000:10017 USPATFULL

TITLE: Human proteasome subunit proteins

INVENTOR(S): Bandman, Olga, Mountain View, CA, United States

Au-Young, Janice, Berkeley, CA, United States Hillman, Jennifer L., San Jose, CA, United States

Goli, Surya K., Sunnyvale, CA, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

RELATED APPLN. INFO.: Division of Ser. No. US 1996-701935, filed on 23 Aug

1996, now patented, Pat. No. US 5843715

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Kemmerer, Elizabeth ASSISTANT EXAMINER: Romeo, David S.

LEGAL REPRESENTATIVE: Price, Esq., Leanne C. Incyte Pharmaceuticals, Inc.

NUMBER OF CLAIMS: 2
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 14 Drawing Figure(s); 18 Drawing Page(s)

LINE COUNT: 1999

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 12 OF 36 USPATFULL

TI Polynucleotide encoding human G-protein coupled receptor

The invention provides a human G-protein coupled receptor (GRecH) and polynucleotides which identify and encode GRecH. The invention also provides expression vectors, host cells, agonists, antibodies and antagonists. The invention also provides methods for treating disorders associated with expression of GRecH.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 1999:155481 USPATFULL

TITLE: Polynucleotide encoding human G-protein coupled

receptor

Lal, Preeti, Santa Clara, CA, United States uegler, Karl J., Menlo Park, C. United States INVENTOR(S):

Shah, Purvi, Sunnyvale, CA, United States

Corley, Neil C., Mountian View, CA, United States Incyte Pharmaceuticals, Inc., Palo Alto, CA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE

US 5994097 19991130 US 1997-919624 19970828 (8) PATENT INFORMATION: APPLICATION INFO.:

Utility DOCUMENT TYPE: Granted FILE SEGMENT: PRIMARY EXAMINER: Mertz, Prema LEGAL REPRESENTATIVE: Incyte Pharma

Incyte Pharmaceuticals Inc.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

14 Drawing Figure(s); 14 Drawing Page(s) NUMBER OF DRAWINGS:

2384 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 13 OF 36 USPATFULL

TI Human DP1 homolog

The invention provides a human DPl homolog (DPlh)) and polynucleotides AΒ which identify and encode DPlh. The invention also provides expression

vectors, host cells, agonists, antibodies and antagonists. The

invention

also provides methods for treating disorders associated with expression of DPlh.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

1999:117297 USPATFULL ACCESSION NUMBER:

Human DP1 homolog TITLE:

Bandman, Olga, Mountain View, CA, United States INVENTOR(S):

Guegler, Karl J., Menlo Park, CA, United States

Shah, Purvi, Sunnyvale, CA, United States

Petithory, Joanne R., Union City, CA, United States Corley, Neil C., Mountain View, CA, United States Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE

US 5958725 19990928 PATENT INFORMATION: US 1997-865336 19970529 (8) APPLICATION INFO .:

Utility DOCUMENT TYPE: FILE SEGMENT: Granted PRIMARY EXAMINER: Feisee, Lila

Lazar-Wesley, Eliane ASSISTANT EXAMINER:

Incyte Pharmaceuticals, Inc. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

6 Drawing Figure(s); 6 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 14 OF 36 USPATFULL

TIRegulator of cell signaling

The present invention provides a human regulator of G-protein signaling AB(HRGS) and polynucleotides which identify and encode HRGS. The

also provides genetically engineered expression vectors and host cells comprising the nucleic acid sequences encoding HRGS and a method for producing HRGS. The invention also provides for agonists, antibodies,

invention

antagonists specifically binding HRGS, and their se, in the prevention and treatment of liseases associated with express n of HRGS. Additionally, the invention provides for the use of antisense molecules to polynucleotides encoding HRGS for the treatment of diseases associated with the expression of HRGS. The invention also provides diagnostic assays which utilize the polynucleotide, or fragments or the complement thereof, and antibodies specifically binding HRGS.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

1999:113602 USPATFULL ACCESSION NUMBER:

Regulator of cell signaling TITLE:

Hillman, Jennifer L., San Jose, CA, United States INVENTOR(S):

Goli, Surya K., Sunnyvale, CA, United States

Incyte Pharmaceuticals, Inc., Palo Alto, CA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5955314 19990921

APPLICATION INFO.: US 1996-748483 19961108 (8)
DOCUMENT TYPE: Utility
FILE SEGMENT: Granted PRIMARY EXAMINER: Feisee, Lila ASSISTANT EXAMINER: Sun-Hoffman, Lin

LEGAL REPRESENTATIVE: Billings, Lucy J. Incyte Pharmaceuticals, Inc.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 6 Drawing Figure(s); 6 Drawing Page(s)

1967 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 15 OF 36 USPATFULL

TI Ras protein

The invention provides two human Ras proteins, referred to collectively AB as "RAPR" and individually as "RAPR-1" and "RAPR-2", and polynucleotides

which identify and encode RAPR. The invention also provides expression vectors, host cells, agonists, antibodies and antagonists. The invention

also provides methods for preventing and treating disorders associated with expression of RAPR.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

1999:102695 USPATFULL ACCESSION NUMBER:

TITLE: Ras protein

Bandman, Olga, Mountain View, CA, United States INVENTOR(S):

Hillman, Jennifer L., Mountain View, CA, United States

Guegler, Karl J., Menlo Park, CA, United States

Tang, Y. Tom, Sunnyvale, CA, United States

Corley, Neil C., Mountain View, CA, United States

Incyte Pharmaceuticals, Inc., Palo Alto, CA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE

US 5945306 19990831 PATENT INFORMATION: US 1997-991946 19971216 (8) APPLICATION INFO.:

Utility DOCUMENT TYPE: Granted FILE SEGMENT: Lau, Kawai PRIMARY EXAMINER:

Mohan-Peterson, Sheela, Billings, Lucy J. Incyte LEGAL REPRESENTATIVE:

Pharmaceuticals, Inc.

NUMBER OF CLAIMS: 13 EXEMPLARY CLAIM: 1

12 Drawing Figure(s); 12 Drawing age(s) NUMBER OF DRAWINGS:

LINE COUNT:

466

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 16 OF 36 USPATFULL

TI Cyclic nucleotide phosphodiesterases

The invention provides human cyclic nucleotide phosphodiesterases AB (PDE8)

and polynucleotides which identify and encode PDE8. The invention also provides expression vectors, host cells, antibodies, agonists, and antagonists. The invention also provides methods for treating or preventing disorders associated with expression of PDE8.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 1999:89000 USPATFULL

Cyclic nucleotide phosphodiesterases TITLE:

Au-Young, Janice, Berkeley, CA, United States INVENTOR(S): Cocks, Benjamin G., Palo Alto, CA, United States Coleman, Roger, Mountain View, CA, United States Seilhamer, Jeffrey J., Los Altos Hills, CA, United

States

Fisher, Douglas A., Groton, CT, United States

PATENT ASSIGNEE(S): Incyte Pharmaceuticals, Inc., Palo Alto, CA, United

States (U.S. corporation)

NUMBER KIND DATE

19990803 US 5932423 PATENT INFORMATION:

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=> d his

(FILE 'HOME' ENTERED AT 15:06:11 ON 02 NOV 2001)

FILE 'MEDLINE, BIOSIS, TOXLIT, EMBASE, DGENE, USPATFULL, WPIDS, JAPIO, JICST-EPLUS, FROSTI, FSTA, CEN, CEABA-VTB, BIOTECHDS, SCISEARCH, AGRICOLA' ENTERED AT 15:07:01 ON 02 NOV 2001

188451 S FLUORESCEIN ARSENICAL HELIX BINDER OR FLASH L1

L229523 S L1 AND MODIFIED

O S L2 AND TARGET SEQUENCE MOTIF L3

36 S L11 AND AGAROSE

L4467 S L2 AND TARGET SEQUENCE

O S POLYPEPTIDE () METHOD () ISOLATION L5209 S POLYPEPTIDE AND METHOD OF ISOLATED L6

1 S L6 AND L4

139 S L4 AND (POLYPEPTIDE) r_8

Ь9 101 S L8 AND ISOLATION

L1046 S L9 AND ACYLATION

38 S L10 AND ALANINE L11

=> s beta alanine

L12

L1315249 BETA ALANINE

=> s acylation and 113

611 ACYLATION AND L13 L14

=> s l14 and alpha-helical region

1 L14 AND ALPHA-HELICAL REGION L15

=> d l15 ti abs ibib tot

L15 ANSWER 1 OF 1 USPATFULL
TI Tetrahydronaphthalene

TI Tetrahydronaphthalene derivatives

AB The present invention is concerned

The present invention is concerned with tetrahydronaphthalene derivatives which are mimics of domains of peptides or proteins which can interact with other proteins or with DNA or RNA through .alpha.-helical conformation, said tetrahydronaphthalene derivatives having the formulae: ##STR1## are valuable aids in the determination of biologically active peptide sequences and are accordingly so-called "research tools". They are, however, also potentially suitable as medicaments.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 97:56790 USPATFULL

TITLE: Tetrahydronaphthalene derivatives

INVENTOR(S): Abrecht, Christine, Lengnau, Switzerland Muller, Klaus, Munchenstein, Switzerland

Obrecht, Daniel, Basle, Switzerland

Trzeciak, Arnold, Schopfheim, Germany, Federal

Republic

of

PATENT ASSIGNEE(S): Hoffmann-La Roche Inc., Nutley, NJ, United States

(U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5644024 19970701
APPLICATION INFO.: US 1994-292128 19940817 (8)

PRIORITY INFORMATION: CH 1993-DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Bond, Robert T.

LEGAL REPRESENTATIVE: Johnston, George W., Tramaloni, Dennis P., Pokras,

Bruce A.

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 3 Drawing Figure(s); 3 Drawing Page(s)

LINE COUNT: 2621

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> d his

(FILE 'HOME' ENTERED AT 15:06:11 ON 02 NOV 2001)

FILE 'MEDLINE, BIOSIS, TOXLIT, EMBASE, DGENE, USPATFULL, WPIDS, JAPIO, JICST-EPLUS, FROSTI, FSTA, CEN, CEABA-VTB, BIOTECHDS, SCISEARCH, AGRICOLA' ENTERED AT 15:07:01 ON 02 NOV 2001

L1 188451 S FLUORESCEIN ARSENICAL HELIX BINDER OR FLASH

L2 29523 S L1 AND MODIFIED

L3 0 S L2 AND TARGET SEQUENCE MOTIF

L4 467 S L2 AND TARGET SEQUENCE

L5 0 S POLYPEPTIDE () METHOD () ISOLATION L6 209 S POLYPEPTIDE AND METHOD OF ISOLATED

L7 1 S L6 AND L4

L8 139 S L4 AND (POLYPEPTIDE)

L9 101 S L8 AND ISOLATION

L10 46 S L9 AND ACYLATION

L11 38 S L10 AND ALANINE
L12 36 S L11 AM AGAROSE
L13 15249 S BETA ALANINE
L14 611 S ACYLATION AND L13
L15 1 S L14 AND ALPHA-HELICAL REGION

=> s 12 and agarose

L16 921 L2 AND AGAROSE

=> s 116 and N-terminus or C-terminus

8 FILES SEARCHED...

L17 61450 L16 AND N-TERMINUS OR C-TERMINUS

=> s 117 and 114

L18 107 L17 AND L14

=> d 118 ti abs ibib 1-10

L18 ANSWER 1 OF 107 USPATFULL

TI Nucleic acids encoding the C140 receptor

AB Nucleic acid molecules encoding the C140 cell surface receptor have

been

cloned and sequenced. The availability of C140 receptor DNA permits the recombinant production of the C140 receptor which can be produced on

the

surface of a cell, including an oocyte. The nucleic acid molecules are useful in an assay for detecting a substance which affects C140

receptor

activity, either receptor agonists or antagonists. Further, the elucidation of the structure of the C140 receptor permits the design of agonist and antagonist compounds which are useful in such assays. The availability of the C140 receptor also permits production of antibodies specifically immunoreactive with one or more antigenic epitopes of the C140 receptor.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2001:167914 USPATFULL

TITLE:

Nucleic acids encoding the C140 receptor

INVENTOR(S):

Sundelin, Johan, Furulund, Sweden

PATENT ASSIGNEE(S):

Scarborough, Robert M., Belmont, CA, United States Cor Therapeutics Inc., South Francisco, CA, United

States (U.S. corporation)

		NUMBER	KIND	DATE
PATENT INFORMATION:	US	6297026	B1	20011002
APPLICATION INFO.:	US	1995-486673		19950607

RELATED APPLN. INFO.:

Division of Ser. No. US 1995-390301, filed on 25 Jan 1995, now abandoned Continuation-in-part of Ser. No.

(8)

US

1993-97938, filed on 26 Jul 1993, now patented, Pat.

No. US 5629174

DOCUMENT TYPE:
FILE SEGMENT:
PRIMARY EXAMINER:

Utility GRANTED

PRIMARY EXAMINER: Kunz, Gary L. ASSISTANT EXAMINER: Hayes, Robert C

LEGAL REPRESENTATIVE:

Morgan, Lewis & Bockius LLP

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

21

1

NUMBER OF DRAWINGS:

20 Drawing Figure(s); 16 Drawing Page(s)

LINE COUNT: 1363
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 2 OF 107 USPATFULL

TI 4-amino-phenylalanine type compounds which inhibit leukocyte adhesion mediated by VLD-4

mediated by VLA-4

Disclosed are compounds which bind VLA-4. Certain of these compounds also inhibit leukocyte adhesion and, in particular, leukocyte adhesion mediated by VLA-4. Such compounds are useful in the treatment of inflammatory diseases in a mammalian patient, e.g., human, such as asthma, Alzheimer's disease, atherosclerosis, AIDS dementia, diabetes, inflammatory bowel disease, rheumatoid arthritis, tissue

transplantation, tumor metastasis and myocardial ischemia. The compounds

can also be administered for the treatment of inflammatory brain diseases such as multiple sclerosis.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:158276 USPATFULL

TITLE: 4-amino-phenylalanine type compounds which inhibit

leukocyte adhesion mediated by VLA-4

INVENTOR(S): Ashwell, Susan, Plainsboro, NJ, United States

Grant, Francine S., San Francisco, CA, United States Konradi, Andrei W., San Francisco, CA, United States

Kreft, Anthony, Langhorne, PA, United States

Lombardo, Louis John, Relle Mead, NJ, United States Pleiss, Michael A., Sunnyvale, CA, United States Sarantakis, Dimitrios, Newtown, PA, United States Semko, Christopher M., Fremont, CA, United States Thorsett, Eugene D., Moss Beach, CA, United States

PATENT ASSIGNEE(S): Athena Neurosciences, Inc., South San Francisco, CA,

United States (U.S. corporation)

American Home Products Corp., Madison, NJ, United

States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6291453 B1 20010918
APPLICATION INFO.: US 1998-126091 19980730 (9)

NUMBER DATE

PRIORITY INFORMATION: US 1997-112019 19970731 (60)

DOCUMENT TYPE:

FILE SEGMENT:

PRIMARY EXAMINER:

ASSISTANT EXAMINER:

Utility

GRANTED

Kifle, Bruck

Patel, Sudhaker B.

LEGAL REPRESENTATIVE: Burns, Doane, Swecker & Mathis LLP

NUMBER OF CLAIMS: 32
EXEMPLARY CLAIM: 1
LINE COUNT: 4347

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 3 OF 107 USPATFULL

TI DERIVATIVES OF GLP-1 ANALOGS

AB The present invention relates to a pharmaceutical composition comprising

a GLP-1 derivative having a lipophilic substituent; and a surfactant.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:123563 USPATFULL

TITLE: DERIVATIVES OF GLP-1 ANALOGS

INVENTOR(S): KNUDSEN, LISELOTTE BJERRE, VALBY, Denmark

HUUSFELDT, PER OLAF, KOBENHAVN K, Denmark

ARSHOLM, NIELS C., VANLOSE, Denmark
OLSEN, HELLE BIRK, ALLEROD, Denmark
BJORN, SOREN ERIK, LYNGBY, Denmark
PEDERSEN, FREDDY ZIMMERDAHL, VARLOSE, Denmark
MADSEN, KJELD, VARLOSE, Denmark

	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 2001011071	A1	20010802	
APPLICATION INFO.:	US 1999-398111	A 1	19990916	(9)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1999-265141, filed

on 8 Mar 1999, PENDING Continuation-in-part of Ser.

No.

US 1999-258750, filed on 26 Feb 1999, PENDING Continuation-in-part of Ser. No. US 1998-38432, filed on 11 Mar 1998, ABANDONED Continuation-in-part of Ser. No. US 1997-918810, filed on 26 Aug 1997, ABANDONED A 371 of International Ser. No. WO 1997-DK340, filed on

22 Aug 1997, UNKNOWN

	NUMBER	DATE	
PRIORITY INFORMATION:	DK 1996-931	19960830	
	DK 1996-1259	19961108	
	DK 1996-1470	19961220	
	DK 1998-263	19980227	
	DK 1998-264	19980227	
	DK 1998-268	19980227	
	EP 1998-610006	19980313	
	DK 1998-507	19980408	
	DK 1998-272	19980227	
	DK 1998-274	19980227	
	DK 1998-508	19980408	
	DK 1998-509	19980408	
	US 1997-35904	19970124	(60)
	US 1997-36226	19970125	(60)
	us 1997-36255	19970124	(60)
	US 1998-78422	19980318	(60)
	US 1998-82478	19980421	(60)
	US 1998-82479	19980421	(60)
	US 1998-82480	19980421	(60)
	us 1998-82802	19980423	(60)
	US 1998-84357	19980505	(60)
DOCUMENT TYPE	Utility		

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: STEVE T ZELSON, NOVO NORDISK OF NORTH AMERICA INC, 405

LEXINGTON AVENUE, SUITE 6400, NEW YORK, NY, 101746401

NUMBER OF CLAIMS: 238 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 1 Drawing Page(s)

LINE COUNT: 15340

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 4 OF 107 USPATFULL

TI Derivatives of GLP-1 analogs

The present invention relates to GLP-1 derivatives having a lipophilic substituent, pharmaceutical compositions comprising same, and methods of

making an using same. The GLP-1 derivatives of the present invention have a protracted profile of action.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:121450 USPATFULL

TITLE: INVENTOR(S): rivatives of GLP-1 analogs udsen, Liselotte Bjerre, Valby enmark

Huusfeldt, Per Olaf, K.o slashed.benhavn K, Denmark Nielsen, Per Franklin, V.ae butted.rl.o slashed.se,

Denmark

Kaarsholm, Niels C., Vanl.o slashed.se, Denmark Olsen, Helle Birk, Aller.o slashed.d, Denmark

Bj.o slashed.rn, S.o slashed.ren Erik, Lyngby, Denmark

Pedersen, Freddy Zimmerdahl, V.ae butted.rl.o

slashed.se, Denmark

NUMBER

Madsen, Kjeld, V.ae butted.rl.o slashed.se, Denmark

Novo Nordisk A/S, Bagsvaerd, Denmark (non-U.S.

DATE

corporation)

KIND NUMBER DATE B1 20010731 US 6268343 PATENT INFORMATION:

APPLICATION INFO.:

PATENT ASSIGNEE(S):

19990226 (9) US 1999-258750

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1998-38432, filed on 11 Mar 1998, now abandoned Continuation-in-part of Ser. No. US 1997-918810, filed on 26 Aug 1997, now abandoned Continuation-in-part of Ser. No. WO

1997-DK340, filed on 22 Aug 1997

PRIORITY INFORMATION:

	·		
DK	1996-931	19960830	
DK	1996-1259	19961108	
DK	1996-1470	19961220	
DK	1998-263	19980227	
DK	1998-264	19980227	
DK	1998-268	19980227	
DK	1998-272	19980227	
DK	1998-274	19980227	
DK	1998-508	19980408	
DK	1998-509	19980408	
US	1997-35904	19970124	(60)

DOCUMENT TYPE:

Utility GRANTED

FILE SEGMENT:

PRIMARY EXAMINER:

Borin, Michael

LEGAL REPRESENTATIVE:

Zelson, Esq., Steve T., Lambiris, Esq., Elias J.

NUMBER OF CLAIMS:

40

EXEMPLARY CLAIM:

1 Drawing Page(s)

NUMBER OF DRAWINGS: LINE COUNT:

14165

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 5 OF 107 USPATFULL

Conformationally constrained backbone cyclized peptide analogs TI

Novel backbone cyclized peptide analogs are formed by means of bridging AB groups attached via the alpha nitrogens of amino acid derivatives to provide novel non-peptidic linkages. Novel building units disclosed are N.sup..alpha. (.omega.-functionalized) amino acids constructed to include a spacer and a terminal functional group. One or more of these N.sup..alpha. (.omega.-functionalized) amino acids are incorporated

into

a peptide sequence, preferably during solid phase peptide synthesis.

The

reactive terminal functional groups are protected by specific protecting

> groups that can be selectively removed to effect either backbone-to-backbone or backbone-to-side chain cyclizations. The invention is specifically exemplified by backbone cyclized bradykinin antagonists having biological activity. Further embodiments of the

invention are sometostatin analogs having one or two ring structures involving backbon cyclization.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:116981 USPATFULL

TITLE: Conformationally constrained backbone cyclized peptide

analogs

INVENTOR(S): Gilon, Chaim, Jerusalem, Israel

Eren, Doron, Rehovot, Israel Zeltser, Irina, Jerusalem, Israel Seri-Levy, Alon, Jerusalem, Israel Gitan, Gal, Jerusalem, Israel Muller, Dan, Jerusalem, Israel

PATENT ASSIGNEE(S): Yissum Research Development Co. of the Hebrew

University, Jerusalem, Israel (non-U.S. corporation)
Peptor Limited, Rehovot, Israel (non-U.S. corporation)

PATENT INFORMATION: US 6265375 B1 20010724 APPLICATION INFO.: US 1998-120237 19980722 (9)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1995-488159, filed on 7

Jun

1995, now patented, Pat. No. US 5811392

NUMBER DATE

PRIORITY INFORMATION: IL 1994-109943 19940608

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Low, Christopher S. F.

ASSISTANT EXAMINER: Gupta, Anish

LEGAL REPRESENTATIVE: Pennie & Edmonds LLP

NUMBER OF CLAIMS: 17
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 3375

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 6 OF 107 USPATFULL

TI Methods for treatment of multiple sclerosis using peptide analogs of human myelin basic protein

AB The present invention is directed toward peptide analogs of human myelin

basic protein. The peptide analog is at least seven amino acids long

and

INVENTOR(S):

derived from residues 83 to 99 of human myelin basic protein. The analogs are altered from the native sequence at least at positions 91, 95, or 97. Additional alterations may be made at other positions. Pharmaceutical compositions containing these peptide analogs are provided. The peptide analogs are useful for treating multiple sclerosis.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:97421 USPATFULL

TITLE: Methods for treatment of multiple sclerosis using

peptide analogs of human myelin basic protein Gaur, Amitabh, San Diego, CA, United States Conlon, Paul, Solana Beach, CA, United States Ling, Nicholas C., San Diego, CA, United States Staehelin, Theophil, Arlesheim, Switzerland

Crowe, Paul D., Encinitas, CA, United States

PATENT ASSIGNEE(S): Neurocrine Biosciences, Inc., San Diego, CA, United

States (U.S. corporation)

wartis AG, Basel, Switzerland (non-U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 6251396 B1 20010626 APPLICATION INFO.: US 1998-137759 19980820 (9)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1994-342408, filed

on 18 Nov 1994

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Borin, Michael

LEGAL REPRESENTATIVE: Seed Intellectual Property Law Group PLLC

NUMBER OF CLAIMS: 3
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 28 Drawing Figure(s); 15 Drawing Page(s)

LINE COUNT: 1583

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 7 OF 107 USPATFULL

TI COPOLYMERIC, HYDROPHOBICALLY MODIFIED POLYASPARTIC ESTERS HAVING INCREASED MOLECULAR MASS AND THEIR USE

AB The invention describes the preparation of high molecular weight copolymeric polyaspartic esters which have been hydrophobically modified

with alkyl radicals having from 6 to 30 carbon atoms.

Copolymers derived from polyamino acids, in which at least 75 mol % of the units present consist of structural units of the general formulae (I), (II) or (III) ##STR1##

in which the structural elements A are identical or different trifunctional hydrocarbon radicals having 2 carbon atoms of the type (A1) or (A2), where one copolymer consists of at least three units of the formula (I), where

R.sup.1 is as defined for R.sup.2, R.sup.3 or R.sup.4, where

R.sup.2 are one or more radicals from the group of alkali metals, alkaline earth metals, hydrogen or ammonium, [NR.sup.5R.sup.6R.sup.7R.sup.8].sup.+, where R.sup.5 to R.sup.8 independently of one another are hydrogen, alkyl or alkenyl having from 1 to 22 carbon atoms or hydroxyalkyl having from 1 to 22 carbon atoms and from 1 to 6 hydroxyl groups and/or their acylation products containing C.sub.1- to C.sub.22-carboxylic radicals,

R.sup.3 are identical or different, straight-chain or branched, saturated or unsaturated alkyl or alkenyl radicals R.sup.9 having from

to 30 carbon atoms, or radicals of the structure --Y--R.sup.9, where Y is an oligo- or polyoxyalkylene chain having from 1 to 100 oxyalkylene units,

R.sup.4 are identical or different, straight-chain or branched, saturated or unsaturated alkyl or alkenyl radicals having from 1 to 5 carbon atoms, the units of the formula (II) are proteinogenic or nonproteinogenic amino acids and are present in an amount of not more than 20% by weight, and

X in the formula (III) is one or more di- or polyfunctional radicals derived from molecular-mass-increasing agents, in particular a di- or polyhydroxy compound, a di- or polyamino compound, or aminoalcohols, having a linear, branched or cyclic, saturated, unsaturated or aromatic hydrocarbon structure, optionally oxo- or aza-substituted with O or N atoms in the chain,

6

and at least in the case one radical R.sup.1 must ssume the meaning

of R.sup.2 and at least one radical R.sup.1 that of R.sup.3 and at least one radical R.sup.1 that of X.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2001:91625 USPATFULL ACCESSION NUMBER:

TITLE: COPOLYMERIC, HYDROPHOBICALLY MODIFIED POLYASPARTIC

ESTERS HAVING INCREASED MOLECULAR MASS AND THEIR USE

GRUNING, BURGHARD, ESSEN, Germany, Federal Republic of INVENTOR(S): SIMPELKAMP, JORG, ESSEN, Germany, Federal Republic of

WEITEMEYER, CHRISTIAN, ESSEN, Germany, Federal

Republic

of

NUMBER KIND DATE US 2001003776 A1 20010614 PATENT INFORMATION: APPLICATION INFO.: US 1999-312222 A1 19990514 (9)

> NUMBER DATE

DE 1998-19822600 PRIORITY INFORMATION: 19980520

Utility DOCUMENT TYPE: APPLICATION FILE SEGMENT:

LEOPOLD PRESSER, SCULLY SCOTT MURPHY & PRESSER, 400 LEGAL REPRESENTATIVE:

GARDEN CITY PLAZA, GARDEN CITY, NY, 11530

NUMBER OF CLAIMS: 10 1 EXEMPLARY CLAIM: 796 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 8 OF 107 USPATFULL

Cell adhesion ihibiting compounds TI

##STR1## ##STR2##Cyclic peptide of formula (1) where Xaa.sub.1 is ABselected from L-amino acids selected from Phe, Lys and Arg, D-amino acids selected from Phe and Met, the L- and D-amino acid optionally substituted on its .alpha.-carbon or its .alpha.-amino group with a C.sub.1-4 alkyl group; and Melle; Xaa.sub.2, Xaa.sub.3 et Xaa.sub.4 are respectively Leu, Asp and Val, optionally substituted on their .alpha.-carbon or .alpha.-amino group with a C.sub.1-4 alkyl group; X.sup.1 is selected from D-amino acids selected from Ala, Phe, Arg,

Lys,

Trp, hArg(Et).sub.2, Orn(CHMe.sub.2), Orn(Me.sub.2), Lys(CHMe.sub.2)

and

Arg(Pmc), optionally substituted on their .alpha.-carbon or .alpha.-amino group with a C.sub.1-4 alkyl group; Formula (II); NH(CH.sub.2).sub.5 CO; and NH(CH.sub.2).sub.2 S(CH.sub.2).sub.y CO, where y is 1 or 2; X.sup.2 is selected from D-amino acids selected from Ala, Arg, Lys, His, hArg(Et).sub.2, Orm(CHMe.sub.2), and Om(Me.sub.2), optionally substituted on their .alpha.-carbon or .alpha.-amino group with a C.sub.1-4 alkyl group; NH(CH.sub.2)SCH.sub.2 CO; and NH(CH.sub.2).sub.x CO, where x is 2 or 3; Xaa.sub.5 and Xaa.sub.6 are each independently a D-amino acid selected from Ala and Arg, optionally substituted on its .alpha.-carbon or .alpha.-mino group with a

C.sub.1-4

alkyl group; p is 0 or 1; and q is 0 or when p is 1, q is 0 or 1; or a salt thereof. The cyclic peptides inhibit the interaction of vascular cell adhesion molecule-1 and fibronectin with integrin very late antigen

4 (.alpha.4.beta.61) and of mucosal addressin cell adhesion molecule-1 (MAdCAM-1) with integrin .alpha.4.beta.7. They have therapeutic applications such as in rheumatoid arthrids, multiple sclerosis, astlna,

psoriasis, inflarestory bowel disease and insulin-dependent diabetes.

CAS INDEXING IS AVAILABLE FOR THIS PATENT. 2001:75364 USPATFULL ACCESSION NUMBER:

Cell adhesion inibiting compounds TITLE:

INVENTOR(S): Dutta, Anand Swaroop, Macclesfield, United Kingdom Zeneca Limited, London, United Kingdom (non-U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE US 6235711 B1 20010522 US 1998-202831 19981221 (9) PATENT INFORMATION: APPLICATION INFO.:

NUMBER DATE

GB 1996~13112 19960621 PRIORITY INFORMATION:

Utility DOCUMENT TYPE: Granted FILE SEGMENT:

Carlson, Karen Cochrane PRIMARY EXAMINER:

ASSISTANT EXAMINER: Gupta, Anish

LEGAL REPRESENTATIVE: Pillsbury Winthrop LLP

NUMBER OF CLAIMS: 19 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 8 Drawing Figure(s); 5 Drawing Page(s) LINE COUNT: 1825

1825 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 9 OF 107 USPATFULL

Double-stranded peptide nucleic acids \mathtt{TI}

A novel class of compounds, known as peptide nucleic acids, form AB double-stranded structures with one another and with ssDNA. The peptide

nucleic acids generally comprise ligands such as naturally occurring

DNA

bases attached to a peptide backbone through a suitable linker.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:67793 USPATFULL

Double-stranded peptide nucleic acids TITLE:

Norden, Benget, Dorjeskaragatan 15, S-421 60 Vastra INVENTOR(S):

Frolunda, Sweden

Wittung, Pernilla, Djurgardsgatan 27, S-414 62

Gothenburg, Sweden

Buchardt, Ole, Sondergardsvej 73, DK 3500 Vaerlose,

Egholm, Michael, Johnstrup Alle, 3, DK 1923

Fredriksberg, Denmark

Nielsen, Peter E., Hjortevanget 509, DK 2980 Kokkedal,

Denmark

Berg, Rolf, Strandvaenget 6, DK 2960 Rungsted Kyst,

Denmark

NUMBER KIND DATE PATENT INFORMATION: US 6228982 В1 20010508 US 1993-88661 19930702 (8) APPLICATION INFO.:

Continuation-in-part of Ser. No. US 1993-54363, filed RELATED APPLN. INFO.:

on 26 Apr 1993, now patented, Pat. No. US 5539082 Continuation-in-part of Ser. No. WO 1992-EP1219, filed

on 22 May 1992

Utility DOCUMENT TYPE: Granted FILE SEGMENT:

Marschel, Ardin H. PRIMARY EXAMINER:

LEGAL REPRESENTATIVE: Woodcock Washburn Kurtz Mackiewicz & Norris LLP NUMBER OF CLAIMS: EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

GS: 20 Drawing Figure(s); 3 Drawing Page(s)

LINE COUNT: 4722

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L18 ANSWER 10 OF 107 USPATFULL

Methods for treatment of diabetes using peptide analogues of insulin The present invention is directed toward peptide analogues of insulin B chain that are generally derived from peptides comprising residues 9 to 23 of the native B chain sequence. The analogues are altered from the native sequence at position 12, 13, 15 and/or 16, and may be additionally be altered at position 19 and/or other positions. Pharmaceutical compositions containing these peptide analogues are provided. The peptide analogues are useful for treating and inhibiting the development of diabetes.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2001:33422 USPATFULL

TITLE: Methods for treatment of diabetes using peptide

analogues of insulin

INVENTOR(S): Gaur, Amitabh, San Diego, CA, United States

Ling, Nicholas, San Diego, CA, United States

Conlon, Paul J., Solana Beach, CA, United States

PATENT ASSIGNEE(S): Neurocrine Biosciences, San Diego, CA, United States

(U.S. corporation)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1998-28156, filed

on 23 Feb 1998, now abandoned

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Moezie, F. T.

LEGAL REPRESENTATIVE: Seed Intellectual Property Law Group PLLC

NUMBER OF CLAIMS: 2
EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 30 Drawing Figure(s); 15 Drawing Page(s)

LINE COUNT: 1000

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

=> d his

L3

(FILE 'HOME' ENTERED AT 15:06:11 ON 02 NOV 2001)

FILE 'MEDLINE, BIOSIS, TOXLIT, EMBASE, DGENE, USPATFULL, WPIDS, JAPIO, JICST-EPLUS, FROSTI, FSTA, CEN, CEABA-VTB, BIOTECHDS, SCISEARCH, AGRICOLA' ENTERED AT 15:07:01 ON 02 NOV 2001

L1 188451 S FLUORESCEIN ARSENICAL HELIX BINDER OR FLASH

L2 29523 S L1 AND MODIFIED

0 S L2 AND TARGET SEQUENCE MOTIF

L4 467 S L2 AND TARGET SEQUENCE

L5 0 S POLYPEPTIDE () METHOD () ISOLATION L6 209 S POLYPEPTIDE AND METHOD OF ISOLATED

L7 1 S L6 AND L4

L8 139 S L4 AND (POLYPEPTIDE)

L9 101 S L8 AND ISOLATION

L10 46 S L9 AND ACYLATION L11 38 S L10 AND ALANINE

L12 36 S L11 AND AGAROSE

15249 S BETA ATANINE 611 S ACYLA N AND L13 L13 L14 1 S L14 AND ALPHA-HELICAL REGION L15 L16 921 S L2 AND AGAROSE L17 61450 S L16 AND N-TERMINUS OR C-TERMINUS L18 107 S L17 AND L14 => s 12 and immobilized 968 L2 AND IMMOBILIZED L19 => s 119 and 116

L20 363 L19 AND L16

=> s 120 and 16

L21 1 L20 AND L6

=> d 121 ti abs tot

L21 ANSWER 1 OF 1 USPATFULL

Thermophilic polymerase III holoenzyme TI

The present invention relates to gene and amino acid sequences encoding ABDNA polymerase TTT holoenzyme subunits and structural genes from thermophilic organisms. In particular, the present invention provides DNA polymerase III holoenzyme subunits of T. thermophilus. The present invention also provides antibodies and other reagents useful to identify

DNA polymerase III molecules.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

WEST

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Search Results - Record(s) 1 through 10 of 31 returned.

1. Document ID: US 6303119 B1

L7: Entry 1 of 31

File: USPT

Oct 16, 2001

US-PAT-NO: 6303119

DOCUMENT-IDENTIFIER: US 6303119 B1

TITLE: Personal care compositions containing subtilisin enzymes

bound to water insoluble substrates

DATE-ISSUED: October 16, 2001

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Weisgerber; David John Cincinnati OH Allcock; Andrew Campbell Cincinnati OH

US-CL-CURRENT: 424/94.63; 424/443, 424/94.1, 424/94.3

Full Title Citation Front Review Classification Date Reference Claims KWC Draw. Desc Image

2. Document ID: US 6294392 B1

L7: Entry 2 of 31

File: USPT

Sep 25, 2001

US-PAT-NO: 6294392

DOCUMENT-IDENTIFIER: US 6294392 B1

TITLE: Spatially-encoded analyte detection

DATE-ISSUED: September 25, 2001

INVENTOR-INFORMATION:

NAME CITY

STATE ZIP CODE COUNTRY

Kuhr; Werner G. Oak Hills CA
Singhal; Pankaj Berkeley CA
Brazill; Sara Ann Diamond Bar CA

US-CL-CURRENT: 436/518; 435/6

Full Title Citation Front Review Classification Date Reference Claims KMC Draw. Desc Image

3. Document ID: US 6235881 B1

L7: Entry 3 of 31 File: USPT May 22, 2001

US-PAT-NO: 6235881

DOCUMENT-IDENTIFIER: US 6235881 B1

TITLE: Polypeptides encoded by novel HIV-2 proviruses

DATE-ISSUED: May 22, 2001

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Kraus; Gunter Miami FL
Wong-Staal; Flossie San Diego CA
Talbott; Randy L. Princeton NJ
Poeschla; Eric M. San Diego CA

US-CL-CURRENT: 530/350; 530/387.3, 536/23.72

Full Title Citation Front Review Classification Date Reference

KWWC Draw Desc Image

4. Document ID: US 6166178 A

L7: Entry 4 of 31 File: USPT Dec 26, 2000

US-PAT-NO: 6166178

DOCUMENT-IDENTIFIER: US 6166178 A

TITLE: Telomerase catalytic subunit

DATE-ISSUED: December 26, 2000

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Cech; Thomas R. Boulder CO Lingner; Joachim Boulder CO

US-CL-CURRENT: 530/324; 530/827, 530/828, 536/23.2, 536/23.5

Full Title Citation Front Review Classification Date Reference | KMC Draw, Desc Image

5. Document ID: US 6153430 A

L7: Entry 5 of 31 File: USPT Nov 28, 2000

US-PAT-NO: 6153430

DOCUMENT-IDENTIFIER: US 6153430 A

TITLE: Nucleic acid encoding mesothelin, a differentiation antigen present on mesothelium, mesotheliomas and ovarian cancers

DATE-ISSUED: November 28, 2000

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Pastan; Ira Potomac MD Chang; Kai Silver Spring MD

US-CL-CURRENT: 435/325; 435/69.1, 435/69.3, 530/350, 536/23.1,

536/23.5



KobiC | Draw, Desc | Image |

6. Document ID: US 6107462 A

L7: Entry 6 of 31 File: USPT Aug 22, 2000

US-PAT-NO: 6107462

DOCUMENT-IDENTIFIER: US 6107462 A

TITLE: Genes and proteins controlling cholesterol synthesis

DATE-ISSUED: August 22, 2000

INVENTOR - INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Rine; Jasper D. Moraga CA Hampton; Randolph San Diego CA

US-CL-CURRENT: 530/350; 435/69.1, 536/23.5, 536/23.7

Full Title Citation Front Review Classification Date Reference

KWWC Drawn Desc Image

7. Document ID: US 6087103 A

L7: Entry 7 of 31 File: USPT Jul 11, 2000

US-PAT-NO: 6087103

DOCUMENT-IDENTIFIER: US 6087103 A

TITLE: Tagged ligand arrays for identifying target-ligand

interactions

DATE-ISSUED: July 11, 2000

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Burmer; Glenna C. Seattle WA

US-CL-CURRENT: 435/6; 435/7.1, 530/350, 536/23.1



8. Document ID: US 6083502 A

L7: Entry 8 of 31 File: USPT Jul 4, 2000

US-PAT-NO: 6083502

DOCUMENT-IDENTIFIER: US 6083502 A

TITLE: Mesothelium antigen and methods and kits for targeting

it

DATE-ISSUED: July 4, 2000

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Pastan; Ira Potomac MD Chang; Kai Silver Spring MD

US-CL-CURRENT: 424/178.1; 424/133.1, 424/135.1, 424/136.1, 424/138.1, 424/139.1, 424/181.1, 424/183.1, 435/330, 435/331, 530/387.3, 530/387.9, 530/388.5, 530/388.8, 530/391.3,

 $\frac{530}{391.7}$

Full Title Citation Front Review Classification Date Reference



9. Document ID: US 6025477 A

L7: Entry 9 of 31 File: USPT Feb 15, 2000

US-PAT-NO: 6025477

DOCUMENT-IDENTIFIER: US 6025477 A

TITLE: Atherosclerotic plaque specific antigens, antibodies

thereto, and uses thereof

DATE-ISSUED: February 15, 2000

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Calenoff; Emanuel Chicago IL 60611

US-CL-CURRENT: 530/388.2; 435/332, 530/387.3, 530/391.1



10. Document ID: US 5998149 A

L7: Entry 10 of 31 File: USPT Dec 7, 1999

US-PAT-NO: 5998149

DOCUMENT-IDENTIFIER: US 5998149 A

TITLE: Method of detecting transmissible spongiform

encephalopathies

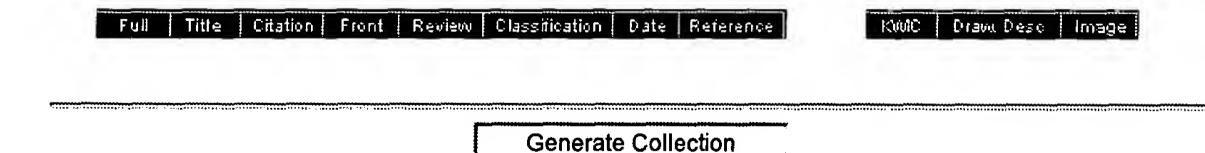
DATE-ISSUED: December 7, 1999

INVENTOR - INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Hsich; Gary Philadelphia PA
Kenney; Kimbra Arlington VA
Gibbs; Clarence J. Washington DC
Harrington; Michael G. La Canada CA

US-CL-CURRENT: 435/7.1; 435/7.92, 435/7.93, 436/149, 436/811



Terms Documents
16 and beads 31

Display 10 Documents, starting with Document: 11

Display Format: CIT Change Format

Generate Collection

L7: Entry 1 of 31

File: USPT

Oct 16, 2001

DOCUMENT-IDENTIFIER: US 6303119 B1

TITLE: Personal care compositions containing subtilisin enzymes

bound to water insoluble substrates

BSPR:

In the medical field, suggestions have been made to diminish the immunogenicity of proteins through yet another method. This method involves attaching unreactive polymers to the protein. U.S. Pat. No. 4,179,337 (Davis, et al.) relates to enzymes coupled to substantially straight chain polyethylene glycol (PEG) or polypropylene glycol (PPG) polymer moieties. While PEG/PPG coupling was found to mitigate the allergenicity of the enzyme, only 15% of the physiological activity was maintained. PCT Application WO 96/17929 (Olsen, et al., published Jun. 13, 1996) relates to the modification of enzymes by conjugating them with suitable polymers. The Olsen application describes modified enzymes which demonstrate a reduction in allergenicity of from 25% to 66% compared to the parent enzyme, while maintaining from 39% to 100% of the activity of the parent.

BSPR:

Another approach to reduce the allergenicity of active proteins has been by granulating, coating or dissolving the active proteins to avoid their becoming airborne. U.S. Pat. No. 4,556,554 (Calvo) discloses cosmetic compositions which comprise enzymes which have been immobilized by attachment to particles of polymeric support. The particles with attached enzymes are dispersed in the cosmetic vehicle. Upon application of the vehicle to the skin, the enzyme is released from the support and is therefore reactivated. Methods such as this address consumer exposure to airborne proteins, however they still leave the substantial risks associated with extended tissue contact with the released enzyme which are deposited on the skin.

BSPR:

Canadian Patent 1,229,808, issued Dec. 1, 1987 teach the immobilization of enzymes, specifically .beta.-galactosidase and .beta.-glucosidase, on cellulosic substrates wherein the enzyme is immobilized by absorption into a agarose gel coating the substrate.

BSPR:

UK Patent Application GB 2,240,040, published Jul. 24, 1991 also teaches immobilized enzymes on substrates. Enzymes, therein as covalently bonded to substrates to provide a

medicated dressing.

BSPR:

The activity of enzymes used in biological equipment such as biosensors, bioseparators, and bioreactors has been enhanced by the use of site-specific attachment of enzymes to equipment surfaces. See Huang et al., "Improving the Activity of Immobilized Subtilisin by Site-specific Attachment to Surface", Analytical Chemistry, 69(22), Nov. 15, 1997. Huang teaches the immobilization of subtilisin enzymes via mutation of serine249 or serine145 to cysteine, and bonding to silica beads functionalized with amino groups.

BSPR:

Nonlimiting examples of synthetic materials useful in the present invention include those selected from the group consisting of acetate fibers, acrylic fibers, cellulose ester fibers, modacrylic fibers, polyamide fibers, polyester fibers, polyolefin fibers, polyvinyl alcohol fibers, rayon fibers, polyurethane foam, and mixtures thereof. Examples of some of these synthetic materials include acrylics such as acrilan, creslan, and the acrylonitrile-based fiber, orlon; cellulose ester fibers such as cellulose acetate, arnel, and acele; polyamides such as nylons (e.g., nylon 6, nylon 66, nylon 610, and the like); polyesters such as fortrel, kodel, and the polyethylene terephthalate fiber, dacron; polyolefins such as polypropylene, polyethylene; polyvinyl acetate fibers; polyurethane foams and mixtures thereof. These and other suitable fibers and the nonwoven materials prepared therefrom are generally described in Riedel, "Nonwoven Bonding Methods and Materials, " Nonwoven World (1987); The Encyclopedia Americana, vol. 11, pp. 147-153, and vol. 26, pp. 566-581 (1984); U.S. Pat. No. 4,891,227, to Thaman et al., issued Jan. 2, 1990; and U.S. Pat. No. 4,891,228 which are all incorporated by reference herein in their entirety.

BSPR:

Nonwoven substrates made from synthetic materials useful in the present invention can also be obtained from a wide variety of commercial sources. Nonlimiting examples of suitable nonwoven layer materials useful herein include HEF 40-047, an apertured hydroentangled material containing about 50% rayon and 50% polyester, and having a basis weight of about 43 grams per square yard (gsy), available from Veratec, Inc., Walpole, Mass.; HEF 140-102, an apertured hydroentangled material containing about 50% rayon and 50% polyester, and having a basis weight of about 56 gsy, available from Veratec, Inc., Walpole, Mass.; Novonet.RTM. 149-616, a thermo-bonded grid patterned material containing about 100% polypropylene, and having a basis weight of about 50 gsy, available from Veratec, Inc., Walpole, Mass.; Novonet.RTM. 149-801, a thermo-bonded grid patterned material containing about 69% rayon, about 25% polypropylene, and about 6% cotton, and having a basis weight of about 75 gsy, available from Veratec, Inc. Walpole, Mass.; Novonet.RTM. 149-191, a thermo-bonded grid patterned material containing about 69% rayon, about 25% polypropylene, and about 6% cotton, and having a basis weight of about 100 gsy,

2 of 4

available from Veratec, Inc. Walpole, Mass.; HEF Nubtex.RTM. 149-801, a nubbed, apertured hydroentangled material, containing about 100% polyester, and having a basis weight of about 70 gsy, available from Veratec, Inc. Walpole, Mass.; Keybak.RTM. 951V, a dry formed apertured material, containing about 75% rayon, about 25% acrylic fibers, and having a basis weight of about 43 gsy, available from Chicopee, New Brunswick, N.J.; Keybak.RTM. 1368, an apertured material, containing about 75% rayon, about 25% polyester, and having a basis weight of about 39 gsy, available from Chicopee, New Brunswick, N.J.; Duralace.RTM. 1236, an apertured, hydroentangled material, containing about 100% rayon, and having a basis weight from about 40 gsy to about 115 gsy, available from Chicopee, New Brunswick, N.J.; Duralace.RTM. 5904, an apertured, hydroentangled material, containing about 100% polyester, and having a basis weight from about 40 gsy to about 115 gsy, available from Chicopee, New Brunswick, N.J.; Sontaro 8868, a hydroentangled material, containing about 50% cellulose and about 50% polyester, and having a basis weight of about 60 gsy, available from Dupont Chemical Corp.

BSPR:

The wipe compositions of the present invention can comprise a wide range of optional ingredients. The CTFA International Cosmetic Ingredient Dictionary, Sixth Edition, 1995, which is incorporated by reference herein in its entirety, describes a wide variety of nonlimiting cosmetic and pharmaceutical ingredients commonly used in the skin care industry, which are suitable for use in the compositions of the present invention. Nonlimiting examples of functional classes of ingredients are described at page 537 of this reference. Examples of these functional classes include: abrasives, anti-acne agents, anticaking agents, anti-microbial agents, antioxidants, binders, biological additives, bulking agents, chelating agents, chemical additives, colorants, cosmetic astringents, cosmetic biocides, denaturants, drug astringents, emulsifiers, external analgesics, film formers, fragrance components, humectants, mildness enhancers (cationic and nonionic polymers, co-surfactants, lipid moisturizers, hydrocarbon oils, silicone oils, waxes), opacifying agents, plasticizers, preservatives, propellants, reducing agents, skin bleaching agents, skin-conditioning agents (emollient, humectants, miscellaneous, and occlusive), skin protectants, solvents, foam boosters, hydrotropes, solubilizing agents, stabilizers, suspending agents, sunscreen agents, surfactants (anionic, cationic, amphoteric, zwitterionic), ultraviolet light absorbers, and viscosity increasing agents (aqueous and nonaqueous). Examples of other functional classes of materials useful herein that are well known to one of ordinary skill in the art include solubilizing agents, sequestrants, and keratolytics, and the like.

DEPR:

A hydroapertured, nonwoven substrate having a basis weight of about 60 gsy comprising 50% rayon and 50% polyester approximately 6 in. by 7.6 in. and a thickness of about 20 mil having a bound active protein per Examples 1-7.



DEPR:

A hydroapertured, nonwoven substrate having a basis weight of about 60 gsy comprising 50% rayon and 50% polyester approximately 6 in. by 7.6 in. and a thickness of about 20 mil having a bound active protein per Examples 1-7.

DEPR:

A hydroapertured, nonwoven substrate having a basis weight of about 60 gsy comprising 50% rayon and 50% polyester approximately 6 in. by 7.6 in. and a thickness of about 20 mil having a bound active protein per Examples 1-7.

DEPR:

A hydroapertured, nonwoven substrate having a basis weight of about 60 gsy comprising 50% rayon and 50% polyester approximately 6 in. by 7.6 in. and a thickness of about 20 mil having a bound active protein according to Examples 1-7.

CLPR:

2. A personal care wipe composition according to claim 1 wherein said water insoluble substrate comprises one or more materials selected from the group consisting of silks, keratins, celluloses, acetates, acrylics, cellulose esters, modacrylics, polyamides, polyesters, polyolefins, polyvinyl alcohols, wood pulp, cotton, hemp, jute, flax, acrylics, nylons, polyesters, polyproylenes, polyethylenes, polyvinyl acetates, polyurethanes, rayon, and mixtures thereof.

CLPR:

3. A personal care wipe composition according to claim 2 wherein said water insoluble substrate comprises a nonwoven sheet of fibers selected from the group consisting of rayon fibers, cellulose fibers, polyester fibers, and mixtures thereof.

ORPL:

Cho, M.Y., Einolf, D.M., "Application of <u>Immobilized</u> Cells and Enzymes for Pharmaceutical Production", Pharmaceutical Manufacturing, 1985 (Oct.), 39-42.

ORPL:

Huang, W. et al., "Improving the Activity of Immobilized Subtilisin by Site-specific Attachment to Surfaces", Analytical Chemistry, 1997 (Nov.), vol. 69 (No. 22), 4601-4607.

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Search Results - Record(s) 11 through 20 of 31 returned.

11. Document ID: US 5977322 A

L7: Entry 11 of 31

File: USPT

Nov 2, 1999

US-PAT-NO: 5977322

DOCUMENT-IDENTIFIER: US 5977322 A

TITLE: High affinity human antibodies to tumor antigens

DATE-ISSUED: November 2, 1999

INVENTOR INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Marks; James D. Kensington CA Schier; Robert San Francisco CA

US-CL-CURRENT: 530/388.85; 530/387.3, 530/387.7, 530/388.15, 530/388.22, 530/388.8

Full Title Citation Front Review Classification Date Reference

KWMC | Draw Desc | Image |

12. Document ID: US 5972625 A

L7: Entry 12 of 31

File: USPT

Oct 26, 1999

US-PAT-NO: 5972625

DOCUMENT-IDENTIFIER: US 5972625 A

TITLE: Assays for inhibitors of leukocyte adhesion

DATE-ISSUED: October 26, 1999

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Rosen; Steven D. San Francisco CA Singer; Mark Berkeley CA

Imai; Yasuyuki Tokyo JPX

US-CL-CURRENT: 435/7.2; 435/7.1, 435/7.24, 435/7.71, 435/7.72, 435/7.8, 435/7.9, 435/7.91, 435/7.92, 435/7.93, 435/7.94, 435/7.95

Full Title Citation Front Review Classification Date Reference

KWWC | Draw Desc | Image

13. Document ID: US 5935788 A

L7: Entry 13 of 31

File: USPT

Aug 10, 1999

US-PAT-NO: 5935788

DOCUMENT-IDENTIFIER: US 5935788 A

TITLE: Subtractive hybridization techniques for identifying differentially expressed and commonly expressed nucleic acid

DATE-ISSUED: August 10, 1999

INVENTOR - INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Burmer; Glenna C. Seattle WA Brown; Joseph P. Seattle WA Stewart; Christine C. Seattle WA

US-CL-CURRENT: 435/6; 435/91.2, 536/23.1, 536/24.2, 536/24.33

Full Title Citation Front Review Classification Date Reference

KMMC Draw Desc Image

14. Document ID: US 5883081 A

L7: Entry 14 of 31

File: USPT

Mar 16, 1999

US-PAT-NO: 5883081

DOCUMENT-IDENTIFIER: US 5883081 A

TITLE: Isolation of novel HIV-2 proviruses

DATE-ISSUED: March 16, 1999

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Kraus; Gunter La Jolla CA
Wong-Staal; Flossie San Diego CA
Talbott; Randy Princeton NJ
Poeschla; Eric M. San Diego CA

US-CL-CURRENT: 514/44; 424/160.1, 435/320.1, 435/69.1,

530/388.35, 536/23.1

Full Title Citation Front Review Classification Date Reference

KWIC Draw Desc Image

15. Document ID: US 5753631 A

L7: Entry 15 of 31 File: USPT May 19, 1998

US-PAT-NO: 5753631

DOCUMENT-IDENTIFIER: US 5753631 A

TITLE: Intercellular adhesion mediators

DATE-ISSUED: May 19, 1998

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Paulson; James C. Sherman Oaks CA
Perez; Mary S. Carlsbad CA
Gaeta; Federico C. A. La Jolla CA
Ratcliffe; Robert M. Carlsbad CA

US-CL-CURRENT: 514/25; 514/54, 514/61, 514/62, 514/8, 536/17.2, 536/18.2, 536/18.7, 536/53, 536/54, 536/55, 536/55.1, 536/55.2

Full Title Citation Front Review Classification Date Reference

KWIC Draw, Desc Image

16. Document ID: US 5750332 A

L7: Entry 16 of 31 File: USPT May 12, 1998

US-PAT-NO: 5750332

DOCUMENT-IDENTIFIER: US 5750332 A

TITLE: Peptomers with enhanced immunogenicity

DATE-ISSUED: May 12, 1998

INVENTOR - INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Robey; Frank A. Bethesda MD Harris-Kelson; Tracy A. Mitchellville MD Robert-Guroff; Marjorie Rockville MD

US-CL-CURRENT: 435/5; 435/974, 514/13, 514/2

Full Title Citation Front Review Classification Date Reference

KMMC Draw Desc Image

17. Document ID: US 5726022 A

L7: Entry 17 of 31 File: USPT Mar 10, 1998

DOCUMENT-IDENTIFIER: US 5726022 A

TITLE: Subtractive hybridization and capture methods and kits

for differential isolation of nucleic acids including

disease-associated sequences

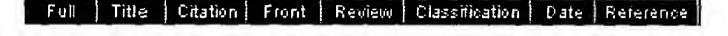
DATE-ISSUED: March 10, 1998

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Burmer; Glenna C. Seattle WA

US-CL-CURRENT: 435/6; 435/91.2, 536/24.2, 536/24.33



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18. Document ID: US 5712103 A

L7: Entry 18 of 31

File: USPT

Jan 27, 1998

US-PAT-NO: 5712103

DOCUMENT-IDENTIFIER: US 5712103 A

TITLE: Diagnostic assay for the prediction of preeclampsia

DATE-ISSUED: January 27, 1998

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Leavitt; John Palo Alto CA
Taylor; Robert N. San Francisco CA
Varma; Madhu Mountain View CA
Shorter; Simon Los Gatos CA

US-CL-CURRENT: 435/7.92; 435/69.6, 435/7.8, 514/3, 514/8

Full Title Citation Front Review Classification Date Reference

KMMC Draw Desc Image

19. Document ID: US 5631133 A

L7: Entry 19 of 31

File: USPT

May 20, 1997

DOCUMENT-IDENTIFIER: US 5631133 A

TITLE: Transition in transcriptional activation by

intracellular hormone receptors at the tumor stage of dermal

fibrosarcoma development

DATE-ISSUED: May 20, 1997

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Hanahan; Douglas San Francisco CA Yamamoto; Keith R. San Francisco CA Vivanco; Maria d. M. San Francisco CA

US-CL-CURRENT: 435/6; 435/69.4



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20. Document ID: US 5604207 A

L7: Entry 20 of 31 File: USPT Feb 18, 1997

US-PAT-NO: 5604207

DOCUMENT-IDENTIFIER: US 5604207 A

TITLE: Sialyl Le.sup.x analogues as inhibitors of cellular

adhesion

DATE-ISSUED: February 18, 1997

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

DeFrees; Shawn A. San Marcos CA Gaeta; Federico C. A. Olivenhain CA Gaudino; John J. Westlake Village CA Zheng; Zhongli Lexington MA

Hayashi; Masaji Kobe JPX

US-CL-CURRENT: 514/25; 514/54, 514/61, 514/62, 536/17.2, 536/55, 536/55.1, 536/55.2, 536/63, 536/64, 536/65

Full Title Citation Front Review Classification Date Reference

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Search Results - Record(s) 21 through 30 of 31 returned.

21. Document ID: US 5559103 A

L7: Entry 21 of 31

File: USPT

Sep 24, 1996

US-PAT-NO: 5559103

DOCUMENT-IDENTIFIER: US 5559103 A

TITLE: Bivalent sialyl X saccharides

DATE-ISSUED: September 24, 1996

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Gaeta; Federico C. A. Foster City CA DeFrees; Shawn A. San Marcos CA

US-CL-CURRENT: 514/54; 514/62, 514/886, 514/887, 530/395, 530/396, 536/53, 536/54, 536/55, 536/55.1, 536/55.2

Full Title Citation Front Review Classification Date Reference

KWIC Draw, Desc Image

22. Document ID: US 5518882 A

L7: Entry 22 of 31

File: USPT

May 21, 1996

US-PAT-NO: 5518882

DOCUMENT-IDENTIFIER: US 5518882 A

TITLE: Immunological methods of component selection and

recovery

DATE-ISSUED: May 21, 1996

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Lund; Garry Edmonton CAX

Wegmann, deceased; late of Edmonton CAX

Mosmann; Timothy Edmonton CAX

US-CL-CURRENT: 435/6; 435/7.21, 435/7.5, 435/7.8, 435/7.93, 436/501, 436/518, 436/541, 436/543

Full Title Citation Front Review Classification Date Reference

KWC Drave Desc Image

23. Document ID: US 5516638 A

L7: Entry 23 of 31

File: USPT

May 14, 1996

US-PAT-NO: 5516638

DOCUMENT-IDENTIFIER: US 5516638 A

TITLE: Immunoassays for the detection of antibodies to Chlamydia trachomatisi in the urine.

DATE-ISSUED: May 14, 1996

INVENTOR - INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Urnovitz; Howard B. San Francisco CA Gottfried; Toby D. Orinda CA Robison; David J. Walnut Creek CA

US-CL-CURRENT: 435/7.32; 435/7.36, 435/7.92, 435/7.93, 435/7.94, 435/7.95, 436/518, 436/530, 436/531, 436/534

Full Title Citation Front Review Classification Date Reference

KMC Draw Desc Image

24. Document ID: US 5424188 A

L7: Entry 24 of 31

File: USPT

Jun 13, 1995

US-PAT-NO: 5424188

DOCUMENT-IDENTIFIER: US 5424188 A

TITLE: Amplified hybridization assay

DATE-ISSUED: June 13, 1995

INVENTOR - INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Schneider; Robert J. New York NY Shenk; Thomas E. Princeton NJ

US-CL-CURRENT: 435/6; 536/24.3

Full Title Citation Front Review Classification Date Reference

KAMC Draw, Desc Image

25. Document ID: US 5318890 A

L7: Entry 25 of 31

File: USPT

Jun 7, 1994

DOCUMENT-IDENTIFIER: US 5318890 A

TITLE: Assays for inhibitors of leukocyte adhesion

DATE-ISSUED: June 7, 1994

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Rosen; Steven San Francisco CA Singer; Mark Berkeley CA Imai; Yasuyuki San Francisco CA Yednock; Ted Fairfax CA

US-CL-CURRENT: 435/7.24; 435/7.1, 435/7.2, 435/7.92, 530/387.3



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26. Document ID: US 5118611 A

L7: Entry 26 of 31

File: USPT

Jun 2, 1992

US-PAT-NO: 5118611

DOCUMENT-IDENTIFIER: US 5118611 A

TITLE: Adenocarcinoma antigen binding methods and reagents

DATE-ISSUED: June 2, 1992

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Smith; Lloyd H. Davis CA Teng; Nelson N. H. Hillsborough CA

US-CL-CURRENT: 435/7.23; 435/344.1, 435/965, 436/548, 436/64, 436/813, 530/387.2, 530/388.15, 530/388.85, 530/808, 530/809, 530/865



KMMC | Draww Desc | Image

27. Document ID: US 5084379 A

L7: Entry 27 of 31

File: USPT

Jan 28, 1992

DOCUMENT-IDENTIFIER: US 5084379 A

TITLE: Fluorometric assay of chymopapain hypersensitivity and

reagents therefor

DATE-ISSUED: January 28, 1992

INVENTOR - INFORMATION:

CITY NAME STATE ZIP CODE COUNTRY

Calenoff; Emanuel Burlingame CA Redwood City CA Jones; Ruth M. Tsay; Yuh-Geng San Jose CA Beigler; Myron A. Los Altos Hills CA

US-CL-CURRENT: 435/7.1; 435/23, 435/24, 435/7.4, 436/513

Full Title Citation Front Review Classification Date Reference

KNAC | Draw Desc | Image

28. Document ID: US 5059654 A

L7: Entry 28 of 31 File: USPT Oct 22, 1991

US-PAT-NO: 5059654

DOCUMENT-IDENTIFIER: US 5059654 A

TITLE: Affinity matrices of modified polysaccharide supports

DATE-ISSUED: October 22, 1991

INVENTOR-INFORMATION:

CITY NAME STATE ZIP CODE COUNTRY

Hou; Kenneth C. Glastonbury CTLiao; Tung-Ping D. Missouri City TXRohan; Robert Columbia CT

US-CL-CURRENT: 525/54.1; 210/198.2, 210/502.1, 210/656, 422/59, 422/70, 422/89, 435/180, 525/54.2, 525/54.21, 530/391.1, 530/391.5, 530/412, 530/413, 536/23.1

Full Title Citation Front Review Classification Date Reference

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29. Document ID: US 4882269 A

L7: Entry 29 of 31 File: USPT Nov 21, 1989

DOCUMENT-IDENTIFIER: US 4882269 A

TITLE: Amplified hybridization assay

DATE-ISSUED: November 21, 1989

INVENTOR-INFORMATION:

STATE ZIP CODE COUNTRY CITY NAME

Schneider; Robert J. New York NY Shenk; Thomas E. Princeton NJ

US-CL-CURRENT: 435/6; 435/18, 435/21, 435/803, 435/810, 436/800, 436/805, 436/808, 536/24.3, 536/24.31, 536/24.32

Full Title Citation Front Review Classification Date Reference

KWIC Draw, Desc Image

30. Document ID: US 4791063 A

L7: Entry 30 of 31

File: USPT

Dec 13, 1988

US-PAT-NO: 4791063

DOCUMENT-IDENTIFIER: US 4791063 A

TITLE: Polyionene transformed modified polysaccharide supports

DATE-ISSUED: December 13, 1988

INVENTOR-INFORMATION:

NAME CITY STATE ZIP CODE COUNTRY

Hou; Kenneth C. S. Glastonbury CTHou; Chung-Jen South Windsor CTChen; Haunn-Lin Vernon

US-CL-CURRENT: 435/243; 435/252.1, 435/308.1, 435/803, 524/27, 524/58, 525/54.3, 526/238.2

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